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OM protein - protein search, using sw model

Run on: March 4, 2004, 15:21:50; Search time 104.702 Seconds

(without alignments)

1397.867 Million cell updates/sec

Title: US-09-668-314C-2

Perfect score: 2687

Sequence: 1 MGALARALLLPLLAQWLLRA.....RPRDPEVVNDESSLVRHRWK 518

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 1586107 seqs, 282547505 residues

Total number of hits satisfying chosen parameters: 1586107

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database: A Geneseq 29Jan04:\*

1: geneseqp1980s:\*

2: geneseqp1990s:\*

3: geneseqp2000s:\*

4: geneseqp2001s:\*

5: genese0qp2002s:\*

6: geneseqp2003as:\*

7: geneseqp2003bs:\*

8: geneseqp2004s:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

### SUMMARIES

Result		% Query					
No.	Score	~ _	Length	DB	ID	Descripti	.on
1	2687	100.0	518	2	AAW61362	Aaw61362	Aspartic
2	2687	100.0	518	2	AAY13799	Aay13799	Human asp
3	2687	100.0	518	2	AAY22239	Aay22239	Human CSP
4	2687	100.0	518	2	AAY41714	Aay41714	Human PRO
5	2687	100.0	518	3	AAY88424	Aay88424	Human asp
6	2687	100.0	518	3	AAB44270	Aab44270	Human PRO
7	2687	100.0	518	4	AAU07201	Aau07201	Human asp
8	2687	100.0	518	4	AAE10628	. Aae10628	Human asp
9	2687	100.0	518	4	AAE10656	Aae10656	Human-Asp

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                                                          Aau29059 Human PRO
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                              AAU29059
14
      2687
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                      518
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16
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                      518
                           5
                              ABB78589
                                                          Abb78589 Human Asp
17
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                              ABU58435
                                                          Abu58435 Human PRO
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43
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45
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## ALIGNMENTS

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     AAW61362;
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DT
     25-MAR-2003
                   (revised)
     25-SEP-1998
                   (first entry)
DT
XX
DE
     Aspartic proteinase ASP1.
XX
KW
     ASP1; Aspartic proteinase; Alzheimer's disease; cancer; melanoma.
XX
OS
     Homo sapiens.
XX
     EP848062-A2.
PN
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XX
PD
    17-JUN-1998.
XX
PF
    01-DEC-1997;
                 97EP-00309648.
XX
                 96GB-00026022.
PR
    14-DEC-1996;
    06-OCT-1997;
                 97US-00999723.
PR
XX
    (SMIK ) SMITHKLINE BEECHAM CORP.
PA
    (SMIK ) SMITHKLINE BEECHAM PLC.
PΑ
XX
                        Chapman CG, Evans JR;
PI
    Powell DJ, Southan C,
XX
DR
    WPI: 1998-314477/28.
    N-PSDB; AAV27962.
DR
XX
    New isolated polynucleotide encodes Aspartic protease polypeptide - used
PT
    to diagnosis, treat and vaccinate against Alzheimer's disease, cancer and
PT
    melanoma.
РΨ
XX
    Claim 11; Page 7; 19pp; English.
PS
XX
    The human ASP1 protein is structurally related to other proteins of the
CC
    Aspartic proteinase family. ASP 1 polypeptides and polynucleotides can be
CC
    used to diagnosis, treat and vaccinate against Alzheimer's disease,
CC
    cancer and melanoma. (Updated on 25-MAR-2003 to correct PR field.)
CC
XX
SO
    Sequence 518 AA;
                      100.0%; Score 2687; DB 2; Length 518;
 Query Match
                      100.0%; Pred. No. 8.6e-231;
 Best Local Similarity
 Matches 518: Conservative
                           0; Mismatches
                                           0: Indels
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Qу
            61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
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Qу
            121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
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Qу
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Qy
            241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
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Qy
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Db
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Qу
            Db
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Qу
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Db
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Qу
            481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Db
RESULT 2
AAY13799
    AAY13799 standard; protein; 518 AA.
XX
AC
    AAY13799;
XX
DT
    21-SEP-1999 (first entry)
XX
DE
    Human aspartyl protease, CSP56.
XX
KW
    CSP56; human; aspartyl protease; diagnosis; neoplasia; tumour;
KW
    breast tumour; colon tumour.
XX
OS
    Homo sapiens.
XX
ΡN
    WO9933963-A1.
XX
PD
    08-JUL-1999.
XX
PF
                  98WO-US026547.
    14-DEC-1998;
XX
PR
    31-DEC-1997;
                  97US-0070112P.
XX
    (CHIR ) CHIRON CORP.
PA
XX
PI
    Giese KW, Xin H;
XX
DR
    WPI; 1999-430240/36.
    N-PSDB; AAX89297.
DR
XX
    Human CSP56 protein for diagnosis of neoplasia.
PT
XX
    Claim 2; Fig 2A; 51pp; English.
PS
XX
    This represents a human CSP56 protein, a novel aspartyl protease. The
CC
CC
    CSP56 protein can be used in methods for diagnosing neoplasia, for
    determining the metastatic potential of a tumour, and for screening test
CC
    compounds for the ability to suppress the metastatic potential of a
CC
    tumour. The tumours are preferably from breast or colon
CC
XX
SO
    Sequence 518 AA;
                       100.0%; Score 2687; DB 2; Length 518;
 Query Match
  Best Local Similarity 100.0%; Pred. No. 8.6e-231;
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Matches
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                                           Indels
                                                     Gaps
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Qу
           Db
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Qγ
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Db
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Qy
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Db
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Qy
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Db
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Qу
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Qу
           Db
       421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Qy
       481 AILLVLIVLLLLPFRCORRPRDPEVVNDESSLVRHRWK 518
           481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Dh
RESULT 3
AAY22239
   AAY22239 standard; protein; 518 AA.
XX
AC
   AAY22239;
XX
DT
   20-SEP-1999
             (first entry)
XX
DE
   Human CSP56, aspartyl-type protease, protein sequence.
XX
KW
   Metastatic marker protein; human; cancer metastasis; breast cancer;
KW
   colon cancer; diagnosis; therapy; tumour; metastatic potential; CSP56;
KW
   aspartyl-type protease.
XX
OS
   Homo sapiens.
XX
PN
   WO9934004-A2.
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XX

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08-JUL-1999.
PD
XX
PF
    24-DEC-1998;
                 98WO-US027608.
XX
    31-DEC-1997;
                 97US-0070112P.
PR
XX
    (CHIR ) CHIRON CORP.
PΑ
XX
PΙ
    Xin H, Giese K;
XX
    WPI: 1999-430248/36.
DR
    N-PSDB; AAX84708.
DR
XX
    New polynucleotides associated with cancer metastasis.
PT
XX
    Claim 4; Page 78-80; 80pp; English.
PS
XX
    This sequence represents a polypeptide of the invention, and is an
CC
    aspartyl-type protease, designated CSP56. The polynucleotides (PNs) of
CC
    the invention encode metastatic marker protein variants. The PNs and
CC
    polypeptides can be used as markers for cancer metastasis. The products
CC
    can be used for identifying metastatic tissue or metastatic potential of
CC
    a tissue, e.g. breast or colon tissue. They can also be used for
CC
    screening test compounds for the ability to suppress the metastatic
CC
    potential of a tumour. The products can be used for developing products
CC
    for the therapy of cancers, particularly breast or colon cancer
CC
XX
SO
    Sequence 518 AA;
                      100.0%; Score 2687; DB 2; Length 518;
  Query Match
                      100.0%; Pred. No. 8.6e-231;
  Best Local Similarity
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  Matches 518; Conservative
                           0; Mismatches
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Qy
            1 MGALARALLIPLIAQWILRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Db
         61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qу
            61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
        121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qy
            121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
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Qу
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DТ
     07-DEC-1999 (first entry)
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    Human PRO852 protein sequence.
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    Human; PRO; EST; expressed sequence tag; PCR primer; hybridisation;
KW
     probe; blood coagulation disorder; cancer; cellular adhesion disorder;
KW
     secreted protein; transmembrane protein.
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XX
OS
     Homo sapiens.
XX
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      15-MAY-1998;
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      15-MAY-1998;
                      98US-0085700P.
PR
      15-MAY-1998;
                      98US-0085704P.
PR
                      98US-0086023P.
PR
      18-MAY-1998;
```

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98US-0086392P.
PR
    22-MAY-1998;
                  98US-0086414P.
PR
    22-MAY-1998;
    22-MAY-1998;
                  98US-0086430P.
PR
    22-MAY-1998;
                  98US-0086486P.
PR
    28-MAY-1998;
                  98US-0087098P.
PR
    28-MAY-1998;
                  98US-0087106P.
PR
    28-MAY-1998;
                  98US-0087208P.
PR
PR
    30-JUL-1998;
                  98US-0094651P.
    11-SEP-1998;
                  98US-0100038P.
PR
XX
    (GETH ) GENENTECH INC.
PΑ
XX
    Wood WI, Goddard A, Gurney A, Yuan J, Baker KP,
PΙ
XX
    WPI; 1999-551358/46.
DR
    N-PSDB; AAZ34056.
DR
XX
    New secreted and transmembrane polypeptides and their polynucleotides,
PΤ
    useful for treating blood coagulation disorders, cancers and cellular
PT
    adhesion disorders.
PT
XX
    Claim 12; Fig 73; 530pp; English.
PS
XX
    The present invention describes secreted and transmembrane polypeptides
CC
    and their polynucleotides. The nucleotide sequences are useful as sources
CC
    of probes, primers, for chromosome mapping, and for generation of
CC
    antisense sequences. They can also be used to create transgenic animals.
CC
    The proteins can be used to treat a variety of diseases and disorders,
CC
    depending on their function. Diseases that may be treated include blood
CC
    coagulation disorders, cancers and cellular adhesion disorders. They may
CC
    also be used to raise antibodies. AAZ33891 to AAZ34338, and AAY41685 to
CC
    AAY41774 represent polynucleotide and polypeptide sequence given in the
CC
CC
    exemplification of the present invention
XX
SQ
    Sequence 518 AA;
                                                  Length 518;
                        100.0%; Score 2687; DB 2;
 Query Match
                       100.0%; Pred. No. 8.6e-231;
 Best Local Similarity
                             0; Mismatches
                                              0;
                                                  Indels
                                                                      0;
 Matches 518; Conservative
           1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qy
             1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Db
          61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qу
             61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
         121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qу
             121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
         181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qy
             181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Db
         241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qу
```

```
241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qy
            301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Db
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
ΟV
            361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Db
        421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Qy
            421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Db
        481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qy
            481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Db
RESULT 5
AAY88424
    AAY88424 standard; protein; 518 AA.
TD
XX
AC
    AAY88424;
XX
DT
    03-AUG-2000 (first entry)
XX
    Human aspartyl protease 1 (Aspl) amino acid sequence.
DE
XX
    Aspartyl protease; aspartase; amyloid precursor protein; APP; Asp 1;
KW
    Alzheimer's disease; beta secretase site.
ΚŴ
XX
os
    Homo sapiens.
XX
PN
    WO200017369-A2.
XX
PD
    30-MAR-2000.
XX
                 99WO-US020881.
PF
    23-SEP-1999;
XX
                 98US-0101594P.
PR
    24-SEP-1998;
XX
    (PHAA ) PHARMACIA & UPJOHN CO.
PA
XX
              Bienkowski MJ, Heinrikson RL, Parodi LA, Yan R;
PΙ
    Gurney ME,
XX
    WPI: 2000-303209/26.
DR
    N-PSDB; AAA15661.
DR
XX
    New enzyme designated human aspartase useful in research into Alzheimer's
PT
    Disease is capable of cleaving amyloid protein precursor at the beta
PT
    secretase site to produce amyloid beta peptide.
PT
XX
    Claim 54; Fig 1; 183pp; English.
PS
XX
    This sequence represents the human aspartyl protease amino acid sequence.
CC
```

CC The invention relates to a protease capable of cleaving the beta secretase site of amyloid precursor protein (APP). The protease contains CC CC a sequence encoding the amino acid sequence DTG and a sequence encoding CC DSG or DTG separated by 100-300 amino acids. When mutated the APP gene CC causes an autosomal dominant form of Alzheimer's disease. APP localises CC to the cell surface membrane and have a single C-terminal transmembrane domain. Proteolytic processing of APP produces the amyloid beta protein, CC which is possibly very important in Alzheimer's disease. The invention CC CC includes a nucleotide sequence encoding the protease, a vector containing CC the nucleotide sequence, and a cell line comprising the vector. Methods CC for screening for inhibitors of beta secretase activity are also given in CC the invention. The human aspartase protein and nucleotide sequences and CC the methods for identifying inhibitors of the protease, are useful in the CC treatment of and research in to Alzheimer's disease XX

SQ Sequence 518 AA;

Query Match 100.0%; Score 2687; DB 3; Length 518; Best Local Similarity 100.0%; Pred. No. 8.6e-231; Matches 518; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

Qу	1	MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL	60
Db	1	MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL	60
Qу	61	ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG	120
Db	61	ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG	120
Qу	121	TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI	180
Db	121	TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI	180
Qу	181	FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA	240
Db	181	FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA	240
Qу	241	GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA	300
Db	241	GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA	300
Qу	301	IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI	360
Db	301		360
Qу	361	YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD	420
Db	361		420
Qу	421	RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG	480
Db	421		480
Qу	481	AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518	
Db	481		

```
RESULT 6
AAB44270
ID
     AAB44270 standard; protein; 518 AA.
XX
AC
     AAB44270;
XX
DT
     08-FEB-2001 (first entry)
XX
DE
     Human PRO852 (UNQ418) protein sequence SEQ ID NO:196.
XX
KW
     Human; secreted protein; transmembrane protein; PRO; EST; cytostatic;
KW
     expressed sequence tag; detection; cancer.
XX
OS
     Homo sapiens.
XX
PN
     WO200053756-A2.
XX
PD
     14-SEP-2000.
XX
PF
     18-FEB-2000; 2000WO-US004341.
XX
     08-MAR-1999;
                    99WO-US005028.
PR
PR
                    99US-0123957P.
     12-MAR-1999;
PR
     29-MAR-1999;
                    99US-0126773P.
PR
     21-APR-1999;
                    99US-0130232P.
     28-APR-1999;
                    99US-0131445P.
PR
PR
     14-MAY-1999;
                    99US-0134287P.
PR
     23-JUN-1999;
                    99US-0141037P.
PR
     26-JUL-1999;
                    99US-0145698P.
PR
     29-OCT-1999;
                    99US-0162506P.
PR
     30-NOV-1999;
                    99WO-US028313.
     02-DEC-1999;
PR
                    99WO-US028551.
PR
     02-DEC-1999;
                    99WO-US028565.
PR
     16-DEC-1999;
                    99WO-US030095.
     30-DEC-1999;
PR
                    99WO-US031243.
PR
     30-DEC-1999;
                    99WO-US031274.
PR
     05-JAN-2000; 2000WO-US000219.
PR
     06-JAN-2000; 2000WO-US000277.
PR
     06-JAN-2000; 2000WO-US000376.
XX
PA
     (GETH ) GENENTECH INC.
XX
PΙ
     Ashkenazi AJ, Baker KP, Botstein D, Desnoyers L,
                                                           Eaton DL;
PΙ
     Ferrara N, Filvaroff E,
                               Fong S, Gao W, Gerber H, Gerritsen ME;
PI
     Goddard A,
                 Godowski PJ,
                               Grimaldi CJ, Gurney AL, Hillan KJ;
PI
     Kljavin IJ, Kuo SS, Napier MA, Pan J, Paoni NF, Roy MA, Shelton DL;
PΙ
     Stewart TA,
                  Tumas D, Williams PM, Wood WI;
XX
DR
     WPI; 2000-611443/58.
DR
     N-PSDB; AAC78500.
XX
PT
     Novel PRO polypeptides and polynucleotides used in detection methods, to
PT
     target bioactive molecules to specific cells, and to modulate cellular
PT
     activities.
XX
PS
     Claim 12; Fig 73; 636pp; English.
```

```
XX
CC
    AAC78458 to AAC78599 represent polynucleotide and EST (expressed sequence
    tag) sequences which encode secreted or transmembrane PRO polypeptides.
CC
    The PRO polynucleotides and polypeptides have cytostatic activity. The
CC
    polynucleotides and polypeptides can be used for detecting the presence
CC
CC
    of PRO polypeptides in samples, for linking bioactive molecules to cells
    and for modulating biological activities of cells, using the polypeptides
CC
    for specific targeting. The polypeptide targeting can be used to kill the
CC
    target cells, e.g. for the treatment of cancers. The polypeptide pairs
CC
CC
    provide specific targeting of bioactive molecules to cells. AAC78600 to
    AAC78987 represent PCR primers and probes used in the isolation of the
CC
CC
    PRO polynucleotide sequences
XX
SQ
    Sequence 518 AA;
 Query Match
                     100.0%; Score 2687; DB 3; Length 518;
 Best Local Similarity
                     100.0%; Pred. No. 8.6e-231;
 Matches 518; Conservative
                           0; Mismatches
                                         0; Indels
                                                        Gaps
                                                               0;
         1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qу
           Db
         1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qy
         61 ALALEPALASPAGAANFLAMVDNLOGDSGRGYYLEMLIGTPPOKLOILVDTGSSNFAVAG 120
           61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
        121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qу
           121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qу
           181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Db
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYOIEILKLEIGGOSLNLDCREYNADKA 300
Qу
           241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qу
           Dh
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qy
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
           Db
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
        421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAOSLSEPILWIVSYALMSVCG 480
Qу
           421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Db
```

481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518

Qу

Db

```
AAU07201
     AAU07201 standard; protein; 518 AA.
ID
XX
     AAU07201;
AC
XX
     24-OCT-2001 (first entry)
DT
XX
     Human aspartyl protease 1 (Asp-1).
DE
XX
     Human; aspartyl protease 1; Asp-1; nootropic; neuroprotective;
KW
ΚW
     aspartyl protease 2; Asp2; amyloid protein precursor; APP;
KW
     beta-secretase; Alzheimer's disease.
XX
OS
     Homo sapiens.
XX
     WO200149097-A2.
PN
XX
     12-JUL-2001.
PD
XX
PF
     09-MAY-2001; 2001WO-IB000797.
XX
     09-MAY-2001; 2001WO-IB000797.
PR
XX
PA
     (BIEN/) BIENKOWSKI M J.
     (GURN/) GURNEY M E.
PA
PΑ
     (HEIN/) HEINRIKSON R L.
PA
     (PARO/) PARODI L A.
     (YANR/) YAN R.
PΑ
XX
     Bienkowski MJ, Gurney ME, Heinrikson RL, Parodi LA, Yan R;
PΙ
XX
DR
     WPI: 2001-502548/55.
     N-PSDB; AAS11701.
DR
XX
     Novel purified polypeptide comprising fragment of mammalian aspartyl
ידים
PT
     protease 2, lacking Asp2 transmembrane domain and retaining beta
РΤ
     secretase activity of Asp2 useful for identifying inhibitors of Asp2
PT
     activity.
XX
PS
     Example 2; Fig 1; 185pp; English.
XX
CC
     The invention relates to a novel purified polypeptide comprising a
CC
     fragment of mammalian aspartyl protease 2 (Asp2) protein which lacks the
CC
     Asp2 transmembrane domain and the Asp2 protein, and where the polypeptide
CC
     and the fragment retain the beta-secretase activity of the mammalian Asp2
CC
     protein. Also included is an isoform of amyloid protein precursor (APP)
CC
     comprising the amino acid sequence of a APP or its fragment containing an
CC
     APP cleavage site recognisable by a mammalian beta-secretase, and further
CC
     comprising two lysine residues at the carboxyl terminus of the amino acid
CC
     sequence of the mammalian APP or APP fragment. The polypeptides are used
CC
     for assaying for modulators of beta-secretase activity; identifying
CC
     agents that inhibit the APP processing activity of human Asp2 aspartyl
CC
     protease (Hu-Asp2); identifying agents that modulate the activity of Asp2
CC
     ; and for reducing cellular production of amyloid beta (Abeta) from APP.
CC
     Agents identified by the above methods are useful for treating
CC
     Alzheimer's disease; and for identifying modulators of amyloid-beta
CC
     (Abeta) peptide production, for use in designing therapeutics for the
```

```
derived from Asp nucleic acid sequences are useful for detecting Hu-Asp
CC
   nucleic acids in in vitro assays and in Northern and Southern blots. The
CC
   present sequence represents the amino acid sequence of human Asp-1
CC
XX
SO
    Sequence 518 AA;
                    100.0%;
                           Score 2687; DB 4; Length 518;
 Query Match
 Best Local Similarity
                    100.0%; Pred. No. 8.6e-231;
 Matches 518; Conservative
                          0; Mismatches
                                       0; Indels
                                                      Gaps
                                                            0;
         1 MGALARALLLPILAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qу
           1 MGALARALLIPLIAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Db
        61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qу
           61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
Qγ
       121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTOGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
           121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
Qy
       181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
           181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Db
       241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qу
           241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYOIEILKLEIGGOSLNLDCREYNADKA 300
Db
       301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qу
           301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Db
       361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Qу
           361 YLRDENSSRSFRITILPOLYIOPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Db
Qу
       421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
           421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Db
Qy
       481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
           Db
        481 AILLVLIVLLLLPFRCORRPRDPEVVNDESSLVRHRWK 518
RESULT 8
AAE10628
ID
   AAE10628 standard; protein; 518 AA.
XX
AC
   AAE10628;
XX
DT
   10-DEC-2001 (first entry)
XX
DE
   Human aspartyl protease 1 (hu-Asp1) protein.
```

treatment or prevention of Alzheimer's disease. Probes and primers

CC

```
XX
KW
     Human; aspartyl protease 1; Asp1; amyloid precursor protein; APP;
KW
     Alzheimer's disease; AD; dementia; neurofibrillary tangle; gliosis;
     amyloid plaque; neuronal loss; proteolytic; nootropic; neuroprotective;
K₩
     chromosome 21.
KW
XX
     Homo sapiens.
OS
XX
FΗ
                     Location/Qualifiers
     Key
     Peptide
FT
                     1. .20
FT
                     /label= Signal peptide
FT
                     21. .518
     Protein
FT
                     /note= "Mature human aspartyl protease 1"
FT
     Domain
                     469. .492
FT
                     /label= Transmembrane domain
XX
PN
     GB2357767-A.
XX
PD
     04-JUL-2001.
XX
     22-SEP-2000; 2000GB-00023315.
PF
XX
PR
     23-SEP-1999;
                    99US-00404133.
PR
     23-SEP-1999;
                    99US-0155493P.
PR
     23-SEP-1999;
                    99WO-US020881.
PR
     13-OCT-1999;
                    99US-00416901.
PR
     06-DEC-1999;
                    99US-0169232P.
XX
     (PHAA ) PHARMACIA & UPJOHN CO.
PA
XX
     Bienkowkski MJ, Gurney M;
PΙ
XX
     WPI; 2001-444208/48.
DR
DR
     N-PSDB; AAD17864.
XX
PT
     Polypeptide comprising fragments of human aspartyl protease with amyloid
PT
     precursor protein processing activity and alpha-secretase activity, for
РΤ
     identifying modulators useful in treating Alzheimer's disease.
XX
PS
     Claim 36; Fig 1; 187pp; English.
XX
CC
     The patent discloses human aspartyl protease 1 (hu-Aspl) or modified Aspl
CC
     proteins which lack transmembrane domain or amino terminal domain or
CC
     cytoplasmic domain and retains alpha-secretase activity and amyloid
CC
     protein precursor (APP) processing activity. The proteins of the
CC
     invention are useful for assaying hu-Aspl alpha-secretase activity, which
CC
     in turn is useful for identifying modulators of hu-Asp1 alpha-secretase
CC
     activity, where modulators that increase hu-Aspl alpha-secretase activity
CC
     are useful for treating Alzheimer's disease (AD) which causes progressive
CC
     dementia with consequent formation of amyloid plaques, neurofibrillary
CC
     tangles, gliosis and neuronal loss. Hu-Aspl protease substrate is useful
CC
     for assaying hu-Aspl proteolytic activity, by contacting hu-Aspl protein
CC
     with the substrate under acidic conditions and determining the level of
CC
     hu-Aspl proteolytic activity. The present sequence is Aspl protein from
CC
     human. Aspl gene is localised on chromosome 21
XX
```

SQ

Sequence 518 AA;

```
100.0%; Score 2687; DB 4; Length 518;
 Query Match
                           Pred. No. 8.6e-231;
                    100.0%;
 Best Local Similarity
                                                            0:
                          0; Mismatches
                                           Indels
                                                      Gaps
                                        0;
 Matches 518; Conservative
         1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qу
           1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Db
        61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qу
           61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
        121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qy
           121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
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Qv
           181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Db
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Qy
           241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Dh
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qy
           301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Db
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Qу
           361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Db
        421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Qу
           421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Db
        481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qу
           481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Db
RESULT 9
AAE10656
    AAE10656 standard; protein; 518 AA.
ID
XX
    AAE10656;
AC
XX
    10-DEC-2001 (first entry)
DT
XX
    Human-Asp 1 protein lacking TM domain and containing (His)6 tag.
DΕ
XX
    Human; aspartyl protease 1; Asp1; amyloid precursor protein; APP;
KW
    Alzheimer's disease; AD; dementia; neurofibrillary tangle; gliosis;
KW
    amyloid plaque; neuronal loss; proteolytic; nootropic; neuroprotective.
KW
XX
    Homo sapiens.
OS
```

```
OS
    Synthetic.
XX
ΡN
    GB2357767-A.
XX
PD
    04-JUL-2001.
XX
PF
    22-SEP-2000; 2000GB-00023315.
XX
                   99US-00404133.
PR
    23-SEP-1999;
                   99US-0155493P.
PR
    23-SEP-1999;
                   99WO-US020881.
PR
    23-SEP-1999;
                   99US-00416901.
PR
    13-OCT-1999;
                   99US-0169232P.
PR
    06-DEC-1999;
XX
     (PHAA ) PHARMACIA & UPJOHN CO.
PΑ
XX
     Bienkowkski MJ, Gurney M;
PΙ
XX
DR
     WPI; 2001-444208/48.
XX
     Polypeptide comprising fragments of human aspartyl protease with amyloid
РΤ
     precursor protein processing activity and alpha-secretase activity, for
PT
     identifying modulators useful in treating Alzheimer's disease.
PT
XX
     Example 14; Page 155-156; 187pp; English.
PS
XX
     The patent discloses human aspartyl protease 1 (hu-Aspl) or modified Aspl
CC
     proteins which lack transmembrane domain or amino terminal domain or
CC
     cytoplasmic domain and retains alpha-secretase activity and amyloid
CC
     protein precursor (APP) processing activity. The proteins of the
CC
     invention are useful for assaying hu-Aspl alpha-secretase activity, which
CC
     in turn is useful for identifying modulators of hu-Asp1 alpha-secretase
CC
     activity, where modulators that increase hu-Asp1 alpha-secretase activity
CC
     are useful for treating Alzheimer's disease (AD) which causes progressive
CC
     dementia with consequent formation of amyloid plaques, neurofibrillary
CC
     tangles, gliosis and neuronal loss. Hu-Aspl protease substrate is useful
CC
     for assaying hu-Asp1 proteolytic activity, by contacting hu-Asp1 protein
CC
     with the substrate under acidic conditions and determining the level of
CC
     hu-Aspl proteolytic activity. The present sequence is human Asp 1 protein
CC
     lacking a transmembrane (TM) domain and containing (His)6 tag. This
CC
     sequence is generated from human Asp 1 protein by the deletion of its C-
CC
     terminal TM domain and addition of hexa-histidine tag at its C-terminus
CC
XX
     Sequence 518 AA;
SQ
                         100.0%; Score 2687; DB 4; Length 518;
  Query Match
                         100.0%; Pred. No. 8.6e-231;
  Best Local Similarity
                                                                   Gaps
                                                                          0;
                                0; Mismatches
                                                 0; Indels
  Matches 518; Conservative
            1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qу
              1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Db
           61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qy
              61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
```

```
121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qу
           121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qу
           181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Db
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qу
           241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
QУ
           301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Db
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Qу
           361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Db
        421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Qу
            421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Db
        481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qу
            481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Db
RESULT 10
AAE06858
    AAE06858 standard; protein; 518 AA.
TD
XX
AC
    AAE06858;
XX
DT
    23-OCT-2001 (first entry)
XX
DE
    Human aspartyl protease 1 (Hu-Aspl) protein.
ХX
    Human; aspartyl protease 1; Asp 1; beta-amyloid precursor protein; APP;
KW
    beta-secretase; Alzheimer's disease; dementia; amyloid plaque; gliosis;
KW
    neurofibrillary tangle; neuronal loss; amyloid-beta peptide; nootropic;
KW
    neuroprotective; antisense therapy; gene therapy; chromosome 21.
KW
XX
OS
    Homo sapiens.
XX
                 Location/Qualifiers
FΗ
    Key
                 1. .20
FT
    Peptide
                 /label= Signal peptide
FT
                 21. .518
FT
    Protein
                 /note= "Mature human aspartyl protease 1 (Hu-Asp1)"
FT
                 469. .492
FT
    Domain
                 /label= Transmembrane domain
FT
XX
    WO200150829-A2.
PN
XX
PD
    19-JUL-2001.
```

```
XX
    09-MAY-2001; 2001WO-IB000799.
PF
XX
    09-MAY-2001; 2001WO-IB000799.
PR
XX
     (BIEN/) BIENKOWSKI M J.
PΑ
     (GURN/) GURNEY M E.
PΑ
PΑ
     (HEIN/) HEINRIKSON R L.
PΑ
     (PARO/) PARODI L A.
     (YANR/) YAN R.
PΑ
XX
    Bienkowski MJ, Gurney ME, Heinrikson RL, Parodi LA, Yan R;
PΙ
XX
DR
    WPI; 2001-483072/52.
DR
    N-PSDB; AAD13020.
XX
    Novel purified polypeptide comprising fragment of mammalian aspartyl
РΤ
    protease 2, lacking Asp2 transmembrane domain and retaining beta
PT
     secretase activity of Asp2 useful for identifying inhibitors of Asp2
PT
РΤ
    activity.
XX
    Example 2; Fig 1; 185pp; English.
PS
XX
    The invention relates to human aspartyl proteases (Hu-Asp), beta-amyloid
CC
     precursor protein (APP) isoforms and their corresponding DNA molecules.
CC
    Human aspartyl proteases can act as beta-secretase proteases useful for
CC
     treating Alzheimer's disease. APP isoforms are useful for identifying
CC
     modulators of amyloid-beta peptide production, for use in designing
CC
     therapeutics for the treatment and prevention of Alzheimer's disease,
CC
     dementia, formation of amyloid plaques, neurofibrillary tangles, gliosis
CC
     and neuronal loss. APP isoforms are also used in methods for identifying
CC
     inhibitors and modulators of human Asp2 activity. The invention relates
CC
     to a method for identifying agents that modulate the activity of human
CC
     aspartyl protease Asp2. Amyloid-beta peptides obtained from APP are used
CC
     as a means to screen in cellular assays for the inhibitors of beta- and
CC
     gamma- secretase. Hu-Asp DNA fragments are useful as probes or primers in
CC
     polymerase chain reactions (PCR). The probes are useful for detecting Hu-
CC
     Asp nucleic acids in in vitro assays and in Northern and Southern blots.
CC
     The present sequence is human aspartyl protease 1 (Hu-Aspl). Hu-Asp 1
CC
     gene is localised on chromosome 21
CC
XX
     Sequence 518 AA;
SO
                        100.0%; Score 2687; DB 4; Length 518;
  Query Match
                        100.0%; Pred. No. 8.6e-231;
  Best Local Similarity
                                                                 Gaps
                                                                         0;
                                                0; Indels
                                                             0;
  Matches 518; Conservative
                               0; Mismatches
           1 MGALARALLIPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qy
             1 MGALARALLIPLIAQWILRAAPELAPAPFTIPLRVAAATNRVVAPTPGPGTPAERHADGI 60
Db
           61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qy
             61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
          121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Οv
```

```
121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qy
           181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Db
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
QУ
            241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qу
           301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Db
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
QУ
            361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Db
        421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Qν
            421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Db
        481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
QУ
            481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Db
RESULT 11
AAE02608
    AAE02608 standard; protein; 518 AA.
TD
XX
AC
    AAE02608;
XX
DT
    10-AUG-2001 (first entry)
XX
    Human Aspartyl protease-1 (Asp-1) deltaTM (His)6 protein.
DE
XX
    Human; alpha-secretase; amyloid precursor protein; APP; therapy;
KW
    Alzheimer's disease; antialzheimer's; aspartyl protease 1; Asp1;
KW
    beta-secretase; Asp-1 deltaTM (His)6 protein.
KW
XX
OS
    Homo sapiens.
    Synthetic.
OS
XX
    WO200123533-A2.
PN
XX
    05-APR-2001.
PD
XX
    22-SEP-2000; 2000WO-US026080.
ΡF
XX
    23-SEP-1999;
                 99US-0155493P.
PR
                 99WO-US020881.
    23-SEP-1999;
PR
                 99US-00416901.
    13-OCT-1999;
PR
                 99US-0169232P.
    06-DEC-1999;
PR
XX
    (PHAA ) PHARMACIA & UPJOHN CO.
PA
XX
```

```
PI
    Gurney M, Bienkowski MJ;
XX
    WPI; 2001-290516/30.
DR
XX
    Enzymes that cleave the alpha-secretase site of the amyloid precursor
РΤ
    protein, useful for the treatment of Alzheimer's disease.
PT
XX
    Example 14; Page 183-184; 189pp; English.
PS
XX
    The present invention relates to enzymes for cleaving the alpha-
CC
    secretase site of the amyloid precursor protein (APP) and methods of
CC
    identifying those enzymes. The methods may be used to identify enzymes
CC
    that may be used to cleave the alpha-secretase cleavage site of the APP
CC
    protein. The enzymes may be used to treat or modulate the progress of
CC
    Alzheimer's disease. The present sequence is human Aspartyl protease-1
CC
    (Asp-1) deltaTM (His)6 protein which is used for the expression of pre-
CC
    pro-human-Aspartyl protease 1 (Asp1). This protein is obtained by
CC
    replacing C-terminal transmembrane and cytoplasmic domains with a
CC
    hexahistidine purification tag in the human Aspartyl protease 1
CC
XX
    Sequence 518 AA;
SO
                     100.0%; Score 2687; DB 4;
 Query Match
                     100.0%; Pred. No. 8.6e-231;
 Best Local Similarity
                                                                0;
                           0; Mismatches
                                          0; Indels
                                                         Gaps
 Matches 518: Conservative
          1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qy
            1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Db
         61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qy
            61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
        121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qу
            121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qу
            181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Db
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qy
            241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qу
            301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Db
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Qy
            361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Db
        421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Qу
            421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Db
```

```
481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qу
              481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Db
RESULT 12
AAE02580
     AAE02580 standard; protein; 518 AA.
ID
XX
     AAE02580;
AC
XX
DT
     10-AUG-2001 (first entry)
XX
     Human aspartyl protease 1 (Asp 1).
DE
XX
     Human; alpha-secretase; amyloid precursor protein; APP; therapy;
KW
     Alzheimer's disease; antialzheimer's; aspartyl protease 1; Asp 1;
KW
KW
     beta-secretase; chromosome 21.
XX
OS
     Homo sapiens.
XX
                     Location/Qualifiers
FH
     Key
                     1. .20
FT
     Peptide
                     /label= Signal peptide
FT
                     22. .62
FT
     Peptide
                     /label= Asp 1 prepropeptide
FT
                     23. .62
FT
     Peptide
                     /label= Asp 1 propeptide
FT
                     63. .518
FT
     Protein
                     /label= Mature human Asp 1_protein
FT
                     /note= "Specifically claimed"
FT
                     87. .89
FT
     Active-site
                     /label= Active_site_1
FT
                     110. .113
FT
     Active-site
                     /label= Active_site_2
FT
     Active-site
                     303. .305
FT
                     /label= Active site 3
FT
                     469. .492
FT
     Domain
                     /label= Transmembrane domain
FT
                      493. .518
FT
     Domain
                      /label= Cytoplasmic domain
FT
                      497. .518
FT
     Region
                      /note= "Peptide #1"
FT
XX
     WO200123533-A2.
PN
XX
     05-APR-2001.
PD
XX
     22-SEP-2000; 2000WO-US026080.
PF
XX
                     99US-0155493P.
     23-SEP-1999;
PR
                     99WO-US020881.
PR
     23-SEP-1999;
     13-OCT-1999;
                     99US-00416901.
PR
                     99US-0169232P.
PR
     06-DEC-1999;
XX
      (PHAA ) PHARMACIA & UPJOHN CO.
PA
```

```
Gurney M,
            Bienkowski MJ;
PΙ
XX
DR
    WPI; 2001-290516/30.
    N-PSDB; AAD06738.
DR
XX
    Enzymes that cleave the alpha-secretase site of the amyloid precursor
PT
    protein, useful for the treatment of Alzheimer's disease.
PΤ
XX
    Claim 29; Fig 1; 189pp; English.
PS
XX
    The present invention relates to enzymes for cleaving the alpha-
CC
    secretase site of the amyloid precursor protein (APP) and methods of
CC
    identifying those enzymes. The methods may be used to identify enzymes
CC
    that may be used to cleave the alpha-secretase cleavage site of the APP
CC
    protein. The enzymes may be used to treat or modulate the progress of
CC
    Alzheimer's disease. The present sequence is human aspartyl protease 1
CC
    (Asp 1). Asp 1 has alpha-secretase protease and beta-secretase protease
CC
    activities. Asp 1 gene is located on chromosome 21
CC
XX
SO
    Sequence 518 AA;
                     100.0%; Score 2687; DB 4; Length 518;
 Query Match
                     100.0%; Pred. No. 8.6e-231;
 Best Local Similarity
                                                                0:
                                                         Gaps
 Matches 518; Conservative
                                          0; Indels
                           0; Mismatches
          1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Ov
           1 MGALARALLIPLIAQWILRAAPELAPAPFTIPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Db
         61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qу
            61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
        121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qy
            121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qу
            181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Db
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qy
            241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qγ
            301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Db
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Qy
            361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Db
        421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Qу
            421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Db
```

ХХ

```
481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qу
              Db
          481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
RESULT 13
AAU29059
     AAU29059 standard; protein; 518 AA.
ID
XX
AC
     AAU29059;
XX
DT
     18-DEC-2001 (first entry)
XX
DE
     Human PRO polypeptide sequence #36.
XX
     PRO polypeptide; mammal; tumour; cancer; human; cattle; horse; sheep;
KW
     dog; cat; pig; goat; rabbit; tumour necrosis factor alpha; TNF-alpha;
KW
     blood; chondrocyte cell; cell proliferation; cell differentiation; colon;
KW
     adrenal; lung; breast; prostate; rectum; cervix; liver; genetic disorder.
ΚW
XX
     Homo sapiens.
OS
XX
PN
     WO200168848-A2.
XX
PD
     20-SEP-2001.
XX
     28-FEB-2001; 2001WO-US006520.
PF
XX
     01-MAR-2000; 2000WO-US005601.
PR
     02-MAR-2000; 2000WO-US005841.
PR
     03-MAR-2000; 2000US-0187202P.
PR
     06-MAR-2000; 2000US-0186968P.
PR
     14-MAR-2000; 2000US-0189320P.
PR
     14-MAR-2000; 2000US-0189328P.
PR
     15-MAR-2000; 2000WO-US006884.
PR
     21-MAR-2000; 2000US-0190828P.
PR
     21-MAR-2000; 2000US-0191007P.
PR
     21-MAR-2000; 2000US-0191048P.
PR
     21-MAR-2000; 2000US-0191314P.
PR
     28-MAR-2000; 2000US-0192655P.
PR
     29-MAR-2000; 2000US-0193032P.
PR
     29-MAR-2000; 2000US-0193053P.
PR
     30-MAR-2000; 2000WO-US008439.
PR
     04-APR-2000; 2000US-0194449P.
PR
     04-APR-2000; 2000US-0194647P.
PR
     11-APR-2000; 2000US-0195975P.
PR
     11-APR-2000; 2000US-0196000P.
PR
     11-APR-2000; 2000US-0196187P.
PR
     11-APR-2000; 2000US-0196690P.
PR
     11-APR-2000; 2000US-0196820P.
PR
     18-APR-2000; 2000US-0198121P.
PR
     18-APR-2000; 2000US-0198585P.
PR
     25-APR-2000; 2000US-0199397P.
PR
     25-APR-2000; 2000US-0199550P.
PR
     25-APR-2000; 2000US-0199654P.
PR
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PR

03-MAY-2000; 2000US-0201516P.

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17-MAY-2000; 2000WO-US013705.
PR
    22-MAY-2000; 2000WO-US014042.
PR
    30-MAY-2000; 2000WO-US014941.
PR
    02-JUN-2000; 2000WO-US015264.
PR
    05-JUN-2000; 2000US-0209832P.
PR
    28-JUL-2000; 2000WO-US020710.
PR
    22-AUG-2000; 2000US-00644848.
PR
    24-AUG-2000; 2000WO-US023328.
PR
    08-NOV-2000; 2000WO-US030952.
PR
    01-DEC-2000; 2000WO-US032678.
PR
    20-DEC-2000; 2000WO-US034956.
PR
XX
     (GETH ) GENENTECH INC.
PA
XX
    Baker KP, Chen J, Desnoyers L, Goddard A, Godowski PJ, Gurney AL;
PΙ
    Pan J, Smith V, Watanabe CK, Wood WI, Zhang Z;
PI
XX
    WPI; 2001-602746/68.
DR
    N-PSDB; AAS45960.
DR
XX
    Novel nucleic acids encoding PRO polypeptides, used to diagnose the
PT
    presence of tumors, such as prostate and breast tumors, in mammals and to
PT
    screen for modulators of the compounds.
PT
XX
    Claim 11; Fig 72; 774pp; English.
PS
XX
     Sequences AAU29024-AAU29328 represent PRO polypeptides of the invention.
CC
     The PRO polypeptides and their associated nucleic acids can be used to
CC
     detect the presence of a tumour in a mammal by comparing the level of
CC
     expression of a PRO polypeptide in a test sample of cells from the animal
CC
     and a control sample of normal cells, whereby a higher level of
CC
     expression in the test sample indicates the presence of a tumour in the
CC
     mammal. Mammals include dogs, cats, cattle, horses, sheep, pigs, goats
CC
     and rabbits but are preferably human. The polypeptides can be used to
CC
     stimulate tumour necrosis factor (TNF) alpha release from human blood,
CC
     when contacted with it. A specific polypeptide can be used to stimulate
CC.
     the proliferation or differentiation of chondrocyte cells. The PRO
CC
     proteins can be used to determine the presence of tumours and also
CC
     susceptibility to tumour development, particularly adrenal, lung, colon,
CC
     breast, prostate, rectal, cervical, or liver tumours, in mammalian
CC
     subjects. The oligonucleotide probes specific for the PRO nucleic acids
CC
     can be used for genetic analysis of individuals with genetic disorders
CC
XX
     Sequence 518 AA;
SO
                         100.0%; Score 2687; DB 4;
  Query Match
                         100.0%; Pred. No. 8.6e-231;
  Best Local Similarity
                                                               0;
                                                                  Gaps
                                                                          0;
                                                 0; Indels
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Qy
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Qy
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Dh
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Qу
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Qу
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Db
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Qy
           301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Db
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Qy
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Db
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Qy
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Db
        481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qу
           481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
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ID
XX
AC
    AAU06602;
XX
    24-OCT-2001 (first entry)
DT
XX
    Human Aspartyl protease 1 (Asp1).
DΕ
XX
    Human; Aspartyl protease; Asp1; Asp2; beta-secretase; nootropic;
KW
    neuroprotective; amyloid protein precursor; APP; Alzheimer's disease;
KW
    amyloid-beta; Abeta.
KW
XX
    Homo sapiens.
OS
XX
PN
    WO200149098-A2.
XX
    12-JUL-2001.
PD
XX
    09-MAY-2001; 2001WO-IB000798.
PF
XX
    09-MAY-2001; 2001WO-IB000798.
PR
XX
    (BIEN/) BIENKOWSKI M J.
PA
    (GURN/) GURNEY M E.
PA
    (HEIN/) HEINRIKSON R L.
PΑ
    (PARO/) PARODI L A.
PΑ
```

```
(YANR/) YAN R.
PA
XX
    Bienkowski MJ, Gurney ME, Heinrikson RL, Parodi LA, Yan R;
ΡI
XX
    WPI; 2001-502549/55.
DR
    N-PSDB; AAS11516.
DR
XX
    Novel purified polypeptide comprising fragment of mammalian aspartyl
PT
    protease 2, lacking Asp2 transmembrane domain and retaining beta
PT
    secretase activity of Asp2 useful for identifying inhibitors of Asp2
PT
PΤ
    activity.
XX
    Example 2; Fig 1; 185pp; English.
PS
XX
    The invention relates to a purified polypeptide comprising a fragment of
CC
    mammalian aspartyl protease (Asp)2 protein which lacks the Asp2
CC
    transmembrane domain and the Asp2 protein, and where the polypeptide and
CC
    the fragment retain the beta-secretase activity of the mammalian Asp2
CC
    protein. The invention also details polynucleotides for the Asp proteins
CC
    and vectors expressing them, and a polypeptide (isoform of amyloid
CC
    protein precursor (APP)) comprising the amino acid sequence of an APP or
CC
    its fragment containing an APP cleavage site recognizable by a mammalian
CC
    beta-secretase, and further comprising two lysine residues at the
CC
    carboxyl terminus of the amino acid sequence of the mammalian APP or APP
CC
    fragment. Also included in the invention are methods of identifying
CC
    modulators or inhibitors of Asp2. Modulators and inhibitors of Asp2 are
CC
    useful for treating Alzheimer's disease. APP is useful in methods for
CC
    identifying inhibitors or modulators of human Asp2 activity and amyloid-
CC
    beta (Abeta) peptide production. APP is also useful in designing
CC
    therapeutics for the treatment or prevention of Alzheimer's disease. APP
CC
    comprising the APP-Sw-beta-secretase peptide sequence (NLDA), which is
CC
    associated with increased levels of Abeta processing is useful in assays
CC
    relating the Alzheimer's research. The expression vector is useful for
CC
     recombinantly expressing APP. Nucleic acids that hybridise to Asp
CC
    oligonucleotides are useful as probes or primers. The probes are useful
CC
     for detecting Hu-Asp nucleic acids in in vitro assays and in Northern and
CC
    Southern blots. The present sequence is human Aspl
CC
XX
SO
     Sequence 518 AA;
                         100.0%; Score 2687; DB 4; Length 518;
  Query Match
                        100.0%; Pred. No. 8.6e-231;
  Best Local Similarity
                                                                         0;
                             0; Mismatches
                                                0; Indels
  Matches 518; Conservative
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Qy
             1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
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Qy
             61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
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Qy
              121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
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Qу
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Qу
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Db
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           421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Db
        481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qy
           481 AILLVLIVLLLLPFRCORRPRDPEVVNDESSLVRHRWK 518
Db
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XX
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XX
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    31-MAY-2002 (first entry)
XX
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XX
    Beta-secretase; enzyme; cleavage site; amyloid protein precursor; APP;
KW
    aspartyl protease; neuroprotective; nootropic; beta-secretase inhibitor;
KW
    Alzheimer's disease.
KW
XX
    Homo sapiens.
OS
XX
    W0200206306-A2.
ΡN
XX
    24-JAN-2002.
ΡD
XX
    19-JUL-2001; 2001WO-US023035.
PF
XX
    19-JUL-2000; 2000US-0219795P.
PR
    12-MAR-2001; 2001US-0275251P.
PR
ΧX
    (PHAA ) PHARMACIA & UPJOHN CO.
PA
XX
                                            Bienkowski MJ;
                                  Emmons TL,
    Yan R, Tomasselli AG,
                        Gurney ME,
PΙ
    Heinrikson RL;
PΙ
XX
DR
    WPI: 2002-216995/27.
XX
    Novel substrates for human aspartyl protease useful for identifying
РΨ
```

modulators of beta secretase activity of aspartyl protease for treating Alzheimer's disease.

Disclosure; Page 161-162; 188pp; English.

The present invention describes an isolated peptide (I) comprising a sequence of at least four amino acids, where the peptide is a substrate for conducting aspartyl protease assays. (I) has neuroprotective and nootropic activities, and can be used as an inhibitor of beta-secretase activity. A beta-secretase modulator from the present invention can be used for inhibiting beta-secretase activity in vivo, and in the manufacture of a medicament for the treatment of Alzheimer's disease. Pharmaceutical compositions from the present invention can be used for treating a disease or condition characterised by an abnormal beta-secretase activity. (I) is useful for identifying agents that modulate the activity of human Asp2 aspartyl protease (Hu-Asp2). (I) is useful as a core structure to construct derivatives. ABL49914 to ABL49925 and ABB06409 to ABB06593 represent sequences used in the exemplification of the present invention

Sequence 518 AA;

PT

PT XX PS

XX

CC

XX SO

100.0%; Score 2687; DB 5; Length 518; Query Match 100.0%; Pred. No. 8.6e-231; Best Local Similarity 0; Matches 518; Conservative 0; Mismatches 0; Indels Gaps 1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60 Qv 1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60 Db 61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120 Qу 61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120 Db 121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180 Qу 121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180 Db 181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240 Qу 181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240 Db 241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300 Qy 241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300 Db 301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360 Qy 301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360 Db 361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420 Qу 361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420 Db 421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480 Qy 421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480 Db

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Db	481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 51	8

Search completed: March 4, 2004, 15:35:42

Job time: 107.702 secs

# GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: March 4, 2004, 15:31:20; Search time 33.6149 Seconds

(without alignments)

795.548 Million cell updates/sec

Title: US-09-668-314C-2

Perfect score: 2687

Sequence: 1 MGALARALLPLLAQWLLRA......RPRDPEVVNDESSLVRHRWK 518

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 389414 seqs, 51625971 residues

Total number of hits satisfying chosen parameters: 389414

Minimum DB seq length: 0

Maximum DB seg length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database: Issued Patents AA:\*

1: /cgn2\_6/ptodata/2/iaa/5A\_COMB.pep:\*

2: /cgn2 6/ptodata/2/iaa/5B\_COMB.pep:\*

3: /cgn2 6/ptodata/2/iaa/6A COMB.pep:\*

4: /cgn2 6/ptodata/2/iaa/6B\_COMB.pep:\*

5: /cgn2 6/ptodata/2/iaa/PCTUS\_COMB.pep:\*

6: /cgn2\_6/ptodata/2/iaa/backfiles1.pep:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

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1	2687	100.0	518	3	US-08-999-723-2	Sequence 2, Appli
2	2687	100.0	518	3	US-09-434-427-2	Sequence 2, Appli
3	2687	100.0	518	4	US-09-548-372D-2	Sequence 2, Appli
4	2687	100.0	518	4	US-09-548-367D-2	Sequence 2, Appli
5	2687	100.0	518	4	US-09-551-853D-2	Sequence 2, Appli
6	2687	100.0	518	4	US-09-215-450-19	Sequence 19, Appl
7	2395	89.1	514	3	US-09-717-432-2	Sequence 2, Appli
8	2395	89.1	514	4	US-09-912-484-2	Sequence 2, Appli
9	1186.5	44.2	501	4	US-09-713-158-2	Sequence 2, Appli
10	1185	44.1	501	4	US-09-548-372D-8	Sequence 8, Appli
11	1185	44.1	501	4	US-09-548-367D-8	Sequence 8, Appli

12	1185	44.1	501	4	US-09-551-853D-8	Sequence	8,	Appli
13	1184.5	44.1	501	4	US-09-724-566A-65	Sequence	65,	Appl
14	1178.5	43.9	501	4	US-09-548-372D-4	Sequence	4,	Appli
15	1178.5	43.9	501	4	US-09-548-367D-4	Sequence		
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19	1175	43.7	503	4	US-09-604-608-3	Sequence		
20	1172.5	43.6	501	4	US-09-009-191-2	Sequence	2,	Appli
21	1171.5	43.6	456	4	US-09-724-566A-43	Sequence		
22	1171.5	43.6	480	4	US-09-724-566A-66	Sequence	66,	Appl
23	1167.5	43.4	444	4	US-09-724-566A-67	Sequence	67,	Appl
24	1160.5	43.2	439	4	US-09-724-566A-69	Sequence	69,	Appl
25	1160.5	43.2	774	4	US-09-009-191-4	Sequence	4,	Appli
26	1139	42.4	452	4	US-09-724-566A-59	Sequence	59,	Appl
27	1139	42.4	453	4	US-09-548-372D-30	Sequence	30,	Appl
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31	1139	42.4	459	4	US-09-548-367D-32	Sequence	32,	Appl
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33	1132	42.1	407	4	US-09-724-566A-58	Sequence	58,	Appl
34	1132	42.1	431	4	US-09-724-566A-74	Sequence	74,	Appl
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42	1127	41.9	459	4	US-09-548-372D-24	Sequence	24,	Appl
43	1127	41.9	459	4	US-09-548-367D-24	Sequence	24,	Appl
44	1127	41.9	459	4	US-09-551-853D-24	Sequence		
45	1124	41.8	420	4	US-09-724-566A-60	Sequence	60,	Appl

## ALIGNMENTS

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RESULT 1
US-08-999-723-2
; Sequence 2, Application US/08999723A
; Patent No. 6025180
; GENERAL INFORMATION:
; APPLICANT: Powell, David J.
; APPLICANT: Southan, Christopher
; APPLICANT: Chapman, Conrad G.
; APPLICANT: Evans, Joanne R.
; TITLE OF INVENTION: ASP1
; FILE REFERENCE: GH70262
; CURRENT APPLICATION NUMBER: US/08/999,723A
; CURRENT FILING DATE: 1997-10-06
; NUMBER OF SEQ ID NOS: 2
; SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 2
    LENGTH: 518
    TYPE: PRT
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Query Match
                   100.0%; Score 2687; DB 3;
                                        Length 518;
 Best Local Similarity
                   100.0%;
                         Pred. No. 2.2e-243:
                        0: Mismatches
                                                   Gaps
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Qy
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Db
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          481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Db
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#### RESULT 2

US-09-434-427-2

- ; Sequence 2, Application US/09434427
- ; Patent No. 6162630
- ; GENERAL INFORMATION:
- ; APPLICANT: POWELL, DAVID J.
- ; APPLICANT: SOUTHAN, CHRISTOPHER
- ; APPLICANT: CHAPMAN, CONRAD G.
- ; APPLICANT: EVANS, JOANNE R.
- ; TITLE OF INVENTION: ASP1
- ; FILE REFERENCE: GH-70262-D1
- ; CURRENT APPLICATION NUMBER: US/09/434,427
- ; CURRENT FILING DATE: 1999-11-04

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EARLIER APPLICATION NUMBER: US 08/999,723
  EARLIER FILING DATE: 1997-10-06
  EARLIER APPLICATION NUMBER: UK 9626022.9
  EARLIER FILING DATE: 1996-12-14
  NUMBER OF SEQ ID NOS: 2
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 SEO ID NO 2
   LENGTH: 518
   TYPE: PRT
   ORGANISM: HOMO SAPIENS
US-09-434-427-2
 Query Match
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                    100.0%; Pred. No. 2.2e-243;
 Best Local Similarity
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                                       0;
 Matches 518; Conservative
                         0; Mismatches
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Qy
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           181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
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       481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
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           481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
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#### RESULT 3

US-09-548-372D-2

<sup>;</sup> Sequence 2, Application US/09548372D

<sup>;</sup> Patent No. 6420534

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; GENERAL INFORMATION:
  APPLICANT: GURNEY ET AL.
  TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR
  TITLE OF INVENTION: THEREOF
  FILE REFERENCE: 29915/6280I
  CURRENT APPLICATION NUMBER: US/09/548,372D
  CURRENT FILING DATE: 2000-04-12
  PRIOR APPLICATION NUMBER: US 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 60/101,594
  PRIOR FILING DATE: 1998-09-24
  NUMBER OF SEQ ID NOS: 73
  SOFTWARE: PatentIn version 3.1
; SEO ID NO 2
   LENGTH: 518
   TYPE: PRT
   ORGANISM: Homo sapiens
US-09-548-372D-2
                     100.0%;
                            Score 2687; DB 4; Length 518;
 Query Match
 Best Local Similarity
                     100.0%; Pred. No. 2.2e-243;
                           0; Mismatches
                                                               0;
                                          0; Indels
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Qу
           61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
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Qy
           121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
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        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qу
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Qу
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Qу
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Db
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Qу
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        481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
           481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
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US-09-548-367D-2
; Sequence 2, Application US/09548367D
; Patent No. 6440698
; GENERAL INFORMATION:
  APPLICANT: GURNEY ET AL.
  TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR
AND USES
  TITLE OF INVENTION: THEREOF
  FILE REFERENCE: 29915/6280H
  CURRENT APPLICATION NUMBER: US/09/548,367D
  CURRENT FILING DATE: 2000-04-12
  PRIOR APPLICATION NUMBER: US 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 60/101,594
  PRIOR FILING DATE: 1998-09-24
  NUMBER OF SEO ID NOS: 73
  SOFTWARE: PatentIn version 3.1
 SEQ ID NO 2
   LENGTH: 518
   TYPE: PRT
   ORGANISM: Homo sapiens
US-09-548-367D-2
                             Score 2687; DB 4;
 Query Match
                      100.0%;
                                              Length 518;
                      100.0%;
                             Pred. No. 2.2e-243;
 Best Local Similarity
 Matches 518; Conservative
                           0; Mismatches
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        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
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RESULT 5
US-09-551-853D-2
; Sequence 2, Application US/09551853D
; Patent No. 6500667
; GENERAL INFORMATION:
  APPLICANT: GURNEY ET AL.
  TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR
AND USES
  TITLE OF INVENTION: THEREOF
  FILE REFERENCE: 29915/6280L
  CURRENT APPLICATION NUMBER: US/09/551,853D
  CURRENT FILING DATE: 2000-04-18
  PRIOR APPLICATION NUMBER: US 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 60/101,594
  PRIOR FILING DATE: 1998-09-24
  NUMBER OF SEQ ID NOS: 73
  SOFTWARE: PatentIn version 3.1
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   LENGTH: 518
   TYPE: PRT
   ORGANISM: Homo sapiens
US-09-551-853D-2
                                              Length 518;
                      100.0%; Score 2687; DB 4;
 Query Match
                             Pred. No. 2.2e-243;
                      100.0%;
 Best Local Similarity
                            0; Mismatches
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                                           0;
                                              Indels
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US-09-215-450-19
; Sequence 19, Application US/09215450
; Patent No. 6635748
; GENERAL INFORMATION:
  APPLICANT: Giese, Klaus
  APPLICANT: Xin, Hong
  TITLE OF INVENTION: METASTATIC BREAST AND COLON CANCER REGULATED GENES
  FILE REFERENCE: 1451.100 / 210030.447
  CURRENT APPLICATION NUMBER: US/09/215,450
  CURRENT FILING DATE: 1998-12-17
  NUMBER OF SEQ ID NOS: 27
  SOFTWARE: FastSEQ for Windows Version 3.0
; SEQ ID NO 19
   LENGTH: 518
   TYPE: PRT
   ORGANISM: human
US-09-215-450-19
                           Score 2687; DB 4;
                                          Length 518;
 Query Match
                    100.0%;
                    100.0%; Pred. No. 2.2e-243;
 Best Local Similarity
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                         0; Mismatches
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                                          Indels
                                                      Gaps
 Matches 518; Conservative
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Qу
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Qу
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           421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
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RESULT 7
US-09-717-432-2
; Sequence 2, Application US/09717432
; Patent No. 6291223
; GENERAL INFORMATION:
  APPLICANT: ZHU, YUAN
  APPLICANT: LI, XIAOTONG
  APPLICANT: CHRISTIE, GARY
            POWELL, DAVID J.
  APPLICANT:
  TITLE OF INVENTION: Mouse Aspartic Secretase-1 (mASP1)
  FILE REFERENCE: GP-70663
  CURRENT APPLICATION NUMBER: US/09/717,432
  CURRENT FILING DATE: 2000-11-21
  PRIOR APPLICATION NUMBER: 60/166,974
  PRIOR FILING DATE: 1999-11-23
  NUMBER OF SEQ ID NOS: 2
  SOFTWARE: FastSEQ for Windows Version 3.0
; SEQ ID NO 2
   LENGTH: 514
   TYPE: PRT
   ORGANISM: MUS MUSCULUS
US-09-717-432-2
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89.1%; Score 2395; DB 3; Length 514;
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 Best Local Similarity
                   88.6%; Pred. No. 5.6e-216;
 Matches 459: Conservative
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        61 ALALEPVRAT----ANFLAMVDNLQGDSGRGYYLEMLIGTPPQKVQILVDTGSSNFAVAG 116
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           117 APHSYIDTYFDSESSSTYHSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNSSFLVNIATI 176
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       181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
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       481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qу
          477 AILLVLILLLLPLHCRHAPRDPEVVNDESSLVRHRWK 514
Db
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### RESULT 8

US-09-912-484-2

- ; Sequence 2, Application US/09912484
- ; Patent No. 6358725
- ; GENERAL INFORMATION:
- ; APPLICANT: Christie, Gary
- ; APPLICANT: Li, Xiaotong
- ; APPLICANT: Powell, David J.
- ; APPLICANT: Zhu, Yuan
- ; TITLE OF INVENTION: Mouse Aspartic Secretase-1 (mASP1)
- ; FILE REFERENCE: GP-70663-D1
- ; CURRENT APPLICATION NUMBER: US/09/912,484
- ; CURRENT FILING DATE: 2001-07-25
- ; PRIOR APPLICATION NUMBER: 60/166,974
- ; PRIOR FILING DATE: 1999-11-23

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PRIOR APPLICATION NUMBER: 09/717,432
  PRIOR FILING DATE: 2000-11-21
  NUMBER OF SEQ ID NOS: 2
  SOFTWARE: FastSEQ for Windows Version 3.0
 SEQ ID NO 2
   LENGTH: 514
   TYPE: PRT
   ORGANISM: MUS MUSCULUS
US-09-912-484-2
                    89.1%; Score 2395; DB 4; Length 514;
 Query Match
                   88.6%; Pred. No. 5.6e-216;
 Best Local Similarity
 Matches 459; Conservative 20; Mismatches
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                                         Indels
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                                                  4; Gaps
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Qy
                     61 ALALEPVRAT----ANFLAMVDNLQGDSGRGYYLEMLIGTPPQKVQILVDTGSSNFAVAG 116
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           117 APHSYIDTYFDSESSSTYHSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNSSFLVNIATI 176
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          177 FESENFFLPGIKWNGILGLAYAALAKPSSSLETFFDSLVAQAKIPDIFSMQMCGAGLPVA 236
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       241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
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#### RESULT 9

US-09-713-158-2

<sup>;</sup> Sequence 2, Application US/09713158

<sup>;</sup> Patent No. 6361975

<sup>;</sup> GENERAL INFORMATION:

<sup>;</sup> APPLICANT: ZHU, YUAN

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APPLICANT: LI, XIAOTONG
  APPLICANT: POWELL, DAVID J.
  APPLICANT: CHRISTIE, GARY
  TITLE OF INVENTION: MOUSE ASPARTIC SECRETASE-2 (MASP-2)
  FILE REFERENCE: GP-70660
  CURRENT APPLICATION NUMBER: US/09/713,158
  CURRENT FILING DATE: 2000-11-15
  PRIOR APPLICATION NUMBER: 60/165,800
  PRIOR FILING DATE: 1999-11-16
  NUMBER OF SEQ ID NOS: 2
  SOFTWARE: FastSEQ for Windows Version 3.0
; SEQ ID NO 2
   LENGTH: 501
   TYPE: PRT
   ORGANISM: MUS MUSCULUS
US-09-713-158-2
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 Matches 238: Conservative 83: Mismatches 167: Indels
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         62 LALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGT 121
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US-09-548-372D-8
; Sequence 8, Application US/09548372D
; Patent No. 6420534
; GENERAL INFORMATION:
  APPLICANT: GURNEY ET AL.
  TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR
AND USES
  TITLE OF INVENTION: THEREOF
  FILE REFERENCE: 29915/6280I
  CURRENT APPLICATION NUMBER: US/09/548,372D
  CURRENT FILING DATE: 2000-04-12
  PRIOR APPLICATION NUMBER: US 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 60/101,594
  PRIOR FILING DATE: 1998-09-24
  NUMBER OF SEQ ID NOS: 73
  SOFTWARE: PatentIn version 3.1
; SEQ ID NO 8
   LENGTH: 501
   TYPE: PRT
   ORGANISM: Mus musculus
US-09-548-372D-8
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 Matches 237; Conservative 83; Mismatches 169; Indels
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; Sequence 8, Application US/09548367D
; Patent No. 6440698
; GENERAL INFORMATION:
; APPLICANT: GURNEY ET AL.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR
AND USES
  TITLE OF INVENTION: THEREOF
  FILE REFERENCE: 29915/6280H
  CURRENT APPLICATION NUMBER: US/09/548,367D
  CURRENT FILING DATE: 2000-04-12
  PRIOR APPLICATION NUMBER: US 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 09/404,133
  PRIOR FILING DATE: 1999-09-23
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  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 60/101,594
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  NUMBER OF SEQ ID NOS: 73
  SOFTWARE: PatentIn version 3.1
; SEQ ID NO 8
   LENGTH: 501
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   ORGANISM: Mus musculus
US-09-548-367D-8
                       44.1%; Score 1185; DB 4; Length 501;
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 Best Local Similarity 46.0%; Pred. No. 1.9e-102;
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; Patent No. 6500667
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; APPLICANT: GURNEY ET AL.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR
AND USES
; TITLE OF INVENTION: THEREOF
; FILE REFERENCE: 29915/6280L
  CURRENT APPLICATION NUMBER: US/09/551,853D
  CURRENT FILING DATE: 2000-04-18
  PRIOR APPLICATION NUMBER: US 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 60/101,594
; PRIOR FILING DATE: 1998-09-24
; NUMBER OF SEQ ID NOS: 73
  SOFTWARE: PatentIn version 3.1
; SEQ ID NO 8
   LENGTH: 501
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   ORGANISM: Mus musculus
US-09-551-853D-8
                     44.1%; Score 1185; DB 4; Length 501;
 Query Match
 Best Local Similarity 46.0%; Pred. No. 1.9e-102;
 Matches 237; Conservative 83; Mismatches 169; Indels 26; Gaps 7;
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; Sequence 65, Application US/09724566A
; Patent No. 6627739
; GENERAL INFORMATION:
  APPLICANT: Anderson, John P.
  APPLICANT: Basi, Guriqbal
  APPLICANT: Doane, Minh Tam
  APPLICANT: Frigon, No. 6627739mand
             John, Varghese
  APPLICANT:
             Power, Michael
  APPLICANT:
             Sinha, Sukanto
  APPLICANT:
  APPLICANT:
            Tatsuno, Gwen
             Tung, Jay
  APPLICANT:
  APPLICANT:
             Wang, Shuwen
             McConlogue, Lisa
  APPLICANT:
  TITLE OF INVENTION: Beta-Secretase Enzyme Compositions and
  TITLE OF INVENTION: Methods
   FILE REFERENCE: 228-US-NEWC2
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CURRENT APPLICATION NUMBER: US/09/724,566A

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CURRENT FILING DATE: 2000-11-28
  PRIOR APPLICATION NUMBER: US 09/501,708
  PRIOR FILING DATE: 2000-02-10
  PRIOR APPLICATION NUMBER: 60/119,571
  PRIOR FILING DATE: 1999-02-10
  PRIOR APPLICATION NUMBER: 60/139,172
  PRIOR FILING DATE: 1999-06-15
  NUMBER OF SEQ ID NOS: 104
  SOFTWARE: FastSEQ for Windows Version 4.0
 SEQ ID NO 65
   LENGTH: 501
   TYPE: PRT
   ORGANISM: Mus musculus
US-09-724-566A-65
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 Best Local Similarity 45.9%; Pred. No. 2.1e-102;
 Matches 237; Conservative 84; Mismatches 170; Indels
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US-09-548-372D-4
; Sequence 4, Application US/09548372D
; Patent No. 6420534
; GENERAL INFORMATION:
  APPLICANT: GURNEY ET AL.
  TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR
AND USES
  TITLE OF INVENTION: THEREOF
  FILE REFERENCE: 29915/6280I
  CURRENT APPLICATION NUMBER: US/09/548,372D
  CURRENT FILING DATE: 2000-04-12
  PRIOR APPLICATION NUMBER: US 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 60/101,594
  PRIOR FILING DATE: 1998-09-24
  NUMBER OF SEQ ID NOS: 73
  SOFTWARE: PatentIn version 3.1
; SEQ ID NO 4
   LENGTH: 501
   TYPE: PRT
   ORGANISM: Homo sapiens
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                      43.9%; Score 1178.5; DB 4; Length 501;
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; Patent No. 6440698
; GENERAL INFORMATION:
; APPLICANT: GURNEY ET AL.
; TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR
AND USES
  TITLE OF INVENTION: THEREOF
 FILE REFERENCE: 29915/6280H
  CURRENT APPLICATION NUMBER: US/09/548,367D
  CURRENT FILING DATE: 2000-04-12
  PRIOR APPLICATION NUMBER: US 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: US 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
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  PRIOR APPLICATION NUMBER: US 60/101,594
  PRIOR FILING DATE: 1998-09-24
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   ORGANISM: Homo sapiens
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Qy
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Search completed: March 4, 2004, 15:42:13 Job time: 34.6149 secs

## GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: March 4, 2004, 15:39:01; Search time 57.8617 Seconds

(without alignments)

1890.324 Million cell updates/sec

Title: US-09-668-314C-2

Perfect score: 2687

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Searched: 809742 segs, 211153259 residues

Total number of hits satisfying chosen parameters: 809742

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Published Applications AA:\* Database :

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/cgn2 6/ptodata/2/pubpaa/PCT NEW PUB.pep:\*

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10: /cgn2\_6/ptodata/2/pubpaa/US09B PUBCOMB.pep:\* /cgn2\_6/ptodata/2/pubpaa/US09C\_PUBCOMB.pep:\* 11:

/cgn2\_6/ptodata/2/pubpaa/US09 NEW PUB.pep:\* 12:

13: /cgn2\_6/ptodata/2/pubpaa/US10A PUBCOMB.pep:\*

14: /cgn2\_6/ptodata/2/pubpaa/US10B\_PUBCOMB.pep:\*

15: /cgn2 6/ptodata/2/pubpaa/US10C PUBCOMB.pep:\*

16: /cgn2 6/ptodata/2/pubpaa/US10 NEW PUB.pep:\*

17: /cgn2 6/ptodata/2/pubpaa/US60 NEW PUB.pep:\*

18: /cgn2 6/ptodata/2/pubpaa/US60 PUBCOMB.pep:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

SUMMARIES

કૃ

Result Query

> No. Score Match Length DB ID

Description

# GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: March 4, 2004, 15:30:05; Search time 28.1043 Seconds

(without alignments)

1772.942 Million cell updates/sec

Title: US-09-668-314C-2

Perfect score: 2687

Sequence: 1 MGALARALLLPLLAQWLLRA.....RPRDPEVVNDESSLVRHRWK 518

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 283366 seqs, 96191526 residues

Total number of hits satisfying chosen parameters: 283366

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database: PIR 78:\*

1: pirl:\*

2: pir2:\*

3: pir3:\*

4: pir4:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

### SUMMARIES

		€				
Result		Query				
No.	Score	Match	Length	DB	ID	Description
1	1178.5	43.9	501	2	A59090	aspartic proteinas
2	367.5	13.7	383	2	JC7573	pepsinogen C - Afr
3	363.5	13.5	377	1	PEMQCJ	gastricsin (EC 3.4
4	355.5	13.2	384	2	A39314	gastricsin (EC 3.4
5	355	13.2	389	2	JE0371	pepsin C (EC 3.4.2
6	353	13.1	388	2	A29937	gastricsin (EC 3.4
7	351.5	13.1	388	2	JC7246	pepsinogen C - com
8	324.5	12.1	394	2	в43356	gastricsin (EC 3.4
9	320	11.9	385	2	JC7575	pepsinogen A - bul
10	320	11.9	402	1	REMSK	renin (EC 3.4.23.1
11	313.5	11.7	509	2	s66516	oryzasin (EC 3.4.2
12	313	11.6	392	1	A24608	gastricsin (EC 3.4
13	310	11.5	383	2	A41443	pepsin (EC 3.4.23.

1.4	308.5	11.5	412	1	KHHUD	cathepsin D (EC 3.
15	306.5	11.4	410	1	KHMSD	cathepsin D (EC 3.
16	305.5	11.4	401	1	REMSS	renin (EC 3.4.23.1
17	305	11.4	384	2	JC7574	pepsinogen A - Afr
18	305	11.4	407	1	KHRTD	cathepsin D (EC 3.
19	302	11.2	405	2	A25379	saccharopepsin (EC
20	301.5	11.2	398	2	S66465	cathepsin E (EC 3.
21	300.5	11.2	387	2	C38302	pepsin (EC 3.4.23.
22	299	11.1	398	2	I51185	cathepsin D (EC 3.
23	298.5	11.1	387	2	D38302	pepsin (EC 3.4.23.
24	298.5	11.1	400	2	I47099	renin (EC 3.4.23.1
25	297	11.1	388	1	PEHU	pepsin A (EC 3.4.2
26	296	11.0	388	2	A30142	pepsin A (EC 3.4.2
27	296	11.0	388	2	B30142	pepsin A (EC 3.4.2
28	294.5	11.0	388	1	S19684	pepsin A (EC 3.4.2
29	292	10.9	506	2	T07915	probable aspartic
30	291	10.8	388	1	S19682	pepsin A (EC 3.4.2
31	291	10.8	402	1	RERTK	renin (EC 3.4.23.1
32	291	10.8	406	1	REHUK	renin (EC 3.4.23.1
33	290.5	10.8	396	2	S36865	cathepsin E (EC 3.
34	289	10.8	387	2	E38302	pepsin (EC 3.4.23.
35	288	10.7	387	2	B38302	pepsin (EC 3.4.23.
36	288	10.7	388	1	PEMQAJ	pepsin A (EC 3.4.2
37	287.5	10.7	632	2	T45858	hypothetical prote
38	287	10.7	391	2	A43356	cathepsin E (EC 3.
39	287	10.7	396	2	A34401	cathepsin E (EC 3.
40	286.5	10.7	334	2	JC4870	pepsin A (EC 3.4.2
41	286	10.6	382	1	PECH	pepsin A (EC 3.4.2
42	286	10.6	388	1	PEMQAR	pepsin A (EC 3.4.2
43	285.5	10.6	387	2	JC7245	pepsinogen A - com
44	285	10.6	396	2	T47207	aspartic proteinas
45	284.5	10.6	386	1	PEPG	pepsin A (EC 3.4.2

### ALIGNMENTS

```
RESULT 1
A59090
aspartic proteinase (EC 3.4.23.-) BACE precursor - human
N;Alternate names: beta-secretase; beta-site APP cleaving enzyme
C;Species: Homo sapiens (man)
C;Date: 29-Oct-1999 #sequence_revision 29-Oct-1999 #text_change 11-May-2000
C;Accession: A59090
R;Vassar, R.; Bennett, B.D.; Babu-Khan, S.; Kahn, S.; Mendiaz, E.A.; Denis, P.;
Teplow, D.B.; Ross, S.; Amarante, P.; Loeloff, R.; Luo, Y.; Fisher, S.; Fuller,
J.; Edenson, S.; Lile, J.; Jarosinski, M.A.; Biere, A.L.; Curran, E.; Burgess,
T.; Louis, J.C.; Collins, F.; Treanor, J.; Rogers, G.; Citron, M.
Science 286, 735-741, 1999
A;Title: beta-Secretase cleavage of Alzheimer's amyloid precursor protein by the transmembrane aspartic protease BACE.
```

A; Reference number: A59090; MUID: 20002972; PMID: 10531052

A; Note: submitted to GenBank, September 1999

A; Status: not compared with conceptual translation

A; Accession: A59090

A; Molecule type: mRNA A; Residues: 1-501 < VAS>

```
A; Cross-references: GB: AF190725; NID: g6118538; PIDN: AAF04142.1; PID: g6118539
C: Genetics:
A:Gene: BACE
C; Superfamily: beta-secretase
C; Keywords: Alzheimer's disease; aspartic proteinase; brain; glycoprotein;
hydrolase; protein digestion; transmembrane protein; zymogen
F;1-21/Domain: signal sequence #status predicted <SIG>
F;22-45/Domain: propeptide #status predicted <PRO>
F;46-501/Product: acid proteinase BACE #status predicted <MAT>
F;461-477/Domain: transmembrane #status predicted <TRN>
F;93,289/Active site: Asp #status predicted
F;153,172,223,354/Binding site: carbohydrate (Asn) (covalent) #status predicted
F:330-380/Disulfide bonds: #status predicted
                       43.9%; Score 1178.5; DB 2; Length 501;
  Query Match
                       46.2%; Pred. No. 3.8e-80;
  Best Local Similarity
 Matches 240; Conservative 82; Mismatches 164; Indels
                                                                     9;
                                                         33; Gaps
           7 ALLLPLLAQWLLRAAPELAPAPFT----LPLRVAAATNRVVAPTPGPGTPAERHADGLA 61
Qу
            | || |:
                           : 11 1
                                      2 AQALPWLLLWM---GAGVLPAHGTQHGIRLPLRSGLG-----GAPL----GLR 42
Db
          62 LALE--PALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVA 119
Qу
                           - 1
          43 LPRETDEEPEEPGRRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFAVG 102
Db
         120 GTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIAT 179
Qу
                                   || :: |: : || ||
         103 AAPHPFLHRYYQRQLSSTYRDLRKGVYVPYTQGKWEGELGTDLVSIPHGPNVTVRANIAA 162
Db
         180 IFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPV 239
Qу
             163 ITESDKFFINGSNWEGILGLAYAEIARPDDSLEPFFDSLVKQTHVPNLFSLQLCGAGFPL 222
Db
         240 AGS---GTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYN 296
Qy
                  1
         223 NQSEVLASVGGSMIIGGIDHSLYTGSLWYTPIRREWYYEVIIVRVEINGQDLKMDCKEYN 282
Db
         297 ADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFP 356
Qу
                                          :| ||| | || ||
                                                            111: 11
              283 YDKSIVDSGTTNLRLPKKVFEAAVKSIKAASSTEKFPDGFWLGEQLVCWQAGTTPWNIFP 342
Db
         357 KISIYLRDENSSRSFRITILPQLYIQPMMGAGLNY-ECYRFGISPSTNALVIGATVMEGF 415
Qу
                                          : :||:| || |:
                                                         1:11:111
              343 VISLYLMGEVTNQSFRITILPQQYLRPVEDVATSQDDCYKFAISQSSTGTVMGAVIMEGF 402
Db
         416 YVIFDRAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYAL 475
Qу
                                                          | ::| :
                                   :: ||| | |:
                                                         1
             11:111:11:11:11
         403 YVVFDRARKRIGFAVSACHVHDEFRTAAVEGPFVTLDMEDCGYNIPQTDESTLMTIAYVM 462
Db
         476 MSVCGAILLVLIVLLLLPFRCQR--RPRDPEVVNDESSL 512
Qy
              ::| |: :: : |:: :|| | | | : : :| | |
         463 AAIC-ALFMLPLCLMVCQWRCLRCLRQQHDDFADDISLL 500
Db
```

```
pepsinogen C - African clawed frog
N; Alternate names: progastricsin
C; Species: Xenopus laevis (African clawed frog)
C;Date: 30-Jun-2001 #sequence revision 30-Jun-2001 #text change 03-Aug-2001
C:Accession: JC7573; PC7118
R; Ikuzawa, M.; Inokuchi, T.; Kobayashi, K.; Yasumasu, S.
J. Biochem. 129, 147-153, 2001
A; Title: Amphibian pepsinogens: Purification and characterization of Xenopus
pepsinogens, and molecular cloning of Xenopus and bullfrog pepsinogens.
A; Reference number: JC7573; MUID:21064922; PMID:11134969
A: Contents: Stomach
A:Accession: JC7573
A: Molecule type: mRNA
A; Residues: 1-383 <IKU>
A; Cross-references: DDBJ: AB045379
A: Accession: PC7118
A; Molecule type: protein
A; Residues: 17-68 < IK2>
C; Comment: This protein is a zymogen for gastric aspartic proteinase, with
pepsin-like activity.
C; Genetics:
A; Gene: PqC
C; Superfamily: pepsin
C; Keywords: stomach; zymogen
                       13.7%; Score 367.5; DB 2; Length 383;
  Query Match
                       28.9%; Pred. No. 9.1e-20;
  Best Local Similarity
 Matches 132; Conservative 70; Mismatches 154; Indels 101; Gaps
                                                                     25;
           1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qу
                                               | | ||: |:: ::| | :1
           1 MKFLILALVCLQLSEGIIR-----VPLKKFKSMREVMRENGIKAPLVDPAT---KYYNQY 52
Db
          61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qу
                                        :1:: 1
          53 ATAYEP----LSNYMDM-----SYYGEISIGTPPQNFLVLFDTGSSNLWVAS 95
Db
         121 TPHSYIDT-----YFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSF 173
Qу
                          1 1:
          96 T---YCQSQACTNHPLFNPSQSSTYSSNQQQFSLQYGTGSLTGILGYDTVTIQ----- 145
Db
         174 LVNIATIFESENFFL----PG-----IKWNGILGLAYATLAKPSSSLETFFDSLVTQANI 224
Qу
                     : | | | | |
                                    :::||||||||
         146 --NVA--ISQQEFGLSETEPGTNFVYAQFDGILGLAYPSIAVGGAT--TVMQGMM-QQNL 198
Db
         225 PN--VFSMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLE 282
Qу
              199 LNQPIFGFYLSGQ-----SSQNGGEVAFGGVDQNYYTGQIYWTPVTSETYWQIGIQGFS 252
Db
         283 IGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQL 342
Qу
                           :|||:||:||:||:||:||:
                                                        :
                                                              :
                    | :
         253 INGQATGW-CSQ--GCQAIVDTGTSLLTAPQSVFSSLIQSIG-----AQQDQNGQYV 301
Db
         343 ACWTNSETPWSYFPKISIYLRDENSSRSFRITILPQLYI-QPMMGAGLNYECYRFGIS-- 399
Qy
                         302 VSCSNIQN----LPTISFTI----SGVSFPLP--PSAYVLQQSSG-----YC-TIGIMPT 345
Db
```

```
400 -- PSTNA---LVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qу
                         ::|
                              : :| ::|
                                             : | | | | :
          346 YLPSONGOPLWILGDVFLREYYSVYDLGNNQVGFATA 382
Db
RESULT 3
PEMQCJ
gastricsin (EC 3.4.23.3) precursor - Japanese macaque (fragment)
N; Alternate names: pepsin C
C; Species: Macaca fuscata (Japanese macaque)
C; Date: 13-Aug-1986 #sequence revision 19-Oct-1995 #text change 18-Jun-1999
C; Accession: S19683; A00986; A22402; S16066
R; Kageyama, T.; Tanabe, K.; Koiwai, O.
Eur. J. Biochem. 202, 205-215, 1991
A; Title: Development-dependent expression of isozymogens of monkey pepsinogens
and structural differences between them.
A; Reference number: S19681; MUID: 92037645; PMID: 1935977
A; Accession: S19683
A; Molecule type: mRNA
A; Residues: 1-377 <KAG>
A;Cross-references: EMBL:X59754; NID:g38072; PIDN:CAA42426.1; PID:g38073
R; Kageyama, T.; Takahashi, K.
J. Biol. Chem. 261, 4406-4419, 1986
A; Title: The complete amino acid sequence of monkey progastricsin.
A; Reference number: A00986; MUID: 86168133; PMID: 3514597
A; Accession: A00986
A; Molecule type: protein
A; Residues: 6-330, 'V', 332-349, 'VY', 350-377 <KA2>
R; Kageyama, T.; Takahashi, K.
J. Biochem. 97, 1235-1246, 1985
A; Title: Monkey pepsinogens and pepsins. VII. Analysis of the activation process
and determination of the NH2-terminal 60-residue sequence of Japanese monkey
progastricsin, and molecular evolution of pepsinogens.
A; Reference number: A22402; MUID: 85289106; PMID: 3928607
A; Accession: A22402
A; Molecule type: protein
A; Residues: 6-65 < KA3>
C; Comment: This enzyme has more restricted specificity than pepsin A.
C; Comment: The enzyme is activated in a two-step process that gives rise to two
end products. The shorter, Ser-gastricsin, is the major product.
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; gastric juice; hydrolase; protein digestion;
F;1-5/Domain: signal sequence (fragment) #status predicted <SIG>
F;6-377/Product: progastricsin #status experimental <ZYM>
F;6-45/Domain: activation peptide #status experimental <APT>
F;46-377/Product: Gly-gastricsin #status experimental <MIN>
F;49-377/Product: Ser-gastricsin #status experimental <MAT>
F;31-32/Cleavage site: Phe-Leu (pepsin) #status experimental
F;45-46/Cleavage site: Phe-Gly (pepsin) #status experimental
F;48-49/Cleavage site: Leu-Ser (pepsin) #status experimental
F;80,265/Active site: Asp #status predicted
F;93-98,256-260,299-332/Disulfide bonds: #status experimental
                           13.5%; Score 363.5; DB 1; Length 377;
  Query Match
                           28.9%; Pred. No. 1.8e-19;
  Best Local Similarity
  Matches 118; Conservative 65; Mismatches 118; Indels 107; Gaps
                                                                              19;
```

```
56 HADGLALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSN 115
Qγ
           44 HFGDLSVSYEP-----MAYMD-----AAYFGEISIGTPPQNFLVLFDTGSSN 85
Db
        116 FAV-----AGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPK 167
Qу
                     86 LWVPSVYCQSQACTSHS----RFNPSESSTYSTNGQTFSLQYGSGSLTGFFGYDTLTV-- 139
Db
        168 GFNTSFLVNIATIFESENFFLPG----IKWNGILGLAYATLAKPSSSLETFFDSLVTQA 222
QУ
                       :::||:|||| |||
        140 ---QSIQVPNQEFGLSEN--EPGTNFVYAQFDGIMGLAYPTLSVDGAT--TAMQGMVQEG 192
Db
        223 NIPN-VFSMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKL 281
Qу
                             : : :||: :
        193 ALTSPIFSVYLSDQ-----QGSSGGAVVFGGVDSSLYTGQIYWAPVTQELYWQIGIEEF 246
Db
        282 EIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQ 341
Οv
             ||||: || :|||:||:||:||:||:||
        247 LIGGQASGW-CSE--GCQAIVDTGTSLLTVPQQYMSALLQA-----TGAQ 288
Db
        342 LACWTNSETPWSYF-----PKISIYLRDENSSRSFRITILPQLYIQPMMGAGLNY 391
Qу
                 289 -----EDEYGQFLVNCNSIQNLPTLTFII-----NGVEFPLPPSSYI-----LNN 328
Db
        392 ECY-RFGISP-----STNALVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qу
             329 NGYCTVGVEPTYLSAQNSQPLWILGDVFLRSYYSVYDLSNNRVGFATA 376
Db
RESULT 4
A39314
gastricsin (EC 3.4.23.3) precursor - bullfrog
C; Species: Rana catesbeiana (bullfrog)
C;Date: 19-Jun-1992 #sequence revision 19-Jun-1992 #text_change 22-Jun-1999
C; Accession: A39314
R; Yakabe, E.; Tanji, M.; Ichinose, M.; Goto, S.; Miki, K.; Kurokawa, K.; Ito,
H.; Kageyama, T.; Takahashi, K.
J. Biol. Chem. 266, 22436-22443, 1991
A; Title: Purification, characterization, and amino acid sequences of pepsinogens
and pepsins from the esophageal mucosa of bullfrog (Rana catesbeiana).
A; Reference number: A39314; MUID: 92042186; PMID: 1939266
A; Accession: A39314
A; Status: preliminary
A; Molecule type: mRNA
A; Residues: 1-384 < YAK>
A; Cross-references: GB: M73750; NID: g213687; PIDN: AAA49530.1; PID: g213688
C: Superfamily: pepsin
C; Keywords: aspartic proteinase; hydrolase; protein digestion
                      13.2%; Score 355.5; DB 2; Length 384;
  Query Match
  Best Local Similarity 26.5%; Pred. No. 7.2e-19;
  Matches 120; Conservative 73; Mismatches 136; Indels 123; Gaps
                                                                 21;
         23 ELAPAPFTLPLRVAAATNRVV-----APTPGPGTPAERHADGLALALEPALASPAGAAN 76
Qу
                  12 QLSEGIIKVPLKKFKSMREVMRDHGIKAPVVDPAT---KYYNNFATAFEP-----LAN 61
Db
```

```
77 FLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGTPHSYIDT-----Y 129
Qγ
                 62 YMDM-----SYYGEISIGTPPONFLVLFDTGSSNLWV---PSTYCQSQACTNHPQ 108
Dh
         130 FDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFESENFFL- 188
Qу
            1: :||:| :::| || || :| || ||
                                               | \cdot | : | \cdot |
         109 FNPSQSSSYSSNQQQFSLQYGTGSLTGILGYDTVQIQ-----NIA--ISQQEFGLS 157
Db
         189 ---PG----IKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPN--VFSMQMCGAGLP 238
Qу
                     :::||||||| ::|: :: | :: | |: |: :|
         158 VTEPGTNFVYAQFDGILGLAYPSIAEGGAT--TVMQGMI-QQNLINQPLFAFYLSG---- 210
Db
         239 VAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNAD 298
Qy
               : ||| : ||:: : | | |::||: | |:|| : ||:
         211 -QQNSQNGGEVAFGGVDQNYYSGQIYWTPVTSETYWQIGIQGFSVNGQATGW-CSQ--GC 266
Db
         299 KAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQ----LACWTNSET 350
Οv
            : |||:||:|| || || || ::::::
         267 QGIVDTGTSLLTAPQSVFSSLMQSI------GAQQDQNGQYAVSCSNIQSL 311
Db
         351 PWSYFP-----KISIYLRDENS---SRSFRITILPQLYIQPMMGAGLNYECYRFGIS 399
Qу
            312 PTISFTISGVSFPLPPSAYVLQQNSGYCTIGIMPTYLPSQNGQPLW----- 357
Db
         400 PSTNALVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qу
               ::| : :| ::| :||1||1||:
Db
         358 -----ILGDVFLRQYYSVYDLGNNQVGFAAA 383
RESULT 5
JE0371
pepsin C (EC 3.4.23.-) precursor - chicken
N; Alternate names: pepsinogen C
C; Species: Gallus gallus (chicken)
C;Date: 23-Jul-1999 #sequence revision 23-Jul-1999 #text change 11-May-2000
C; Accession: JE0371
R; Sakamoto, N.; Saiga, H.; Yasugi, S.
Biochem. Biophys. Res. Commun. 250, 420-424, 1998
A; Title: Analysis of temporal expression pattern and cis-regulatory sequences of
chicken pepsinogen A and C.
A; Reference number: JE0370; MUID:98440813; PMID:9753645
A; Accession: JE0371
A; Status: preliminary
A; Molecule type: mRNA
A; Residues: 1-389 <SAK>
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; hydrolase
                       13.2%; Score 355; DB 2; Length 389;
  Query Match
  Best Local Similarity 28.7%; Pred. No. 7.9e-19;
 Matches 114; Conservative 58; Mismatches 121; Indels 104; Gaps
          75 ANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGT-----PHSYI 126
Qу
            56 SNFATAYEPLANNMDMSYYGEISIGTPPQNFLVLFDTGSSNLWVPSTLCQSQACANHN-- 113
Db
```

```
127 DTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFN----TS 172
QУ
                               111: ::
         114 --EFDPNESSTFSTQDEFFSLQYGSGSLTGIFGFDTVTI-QGISITNQEFGLSETEPGTS 170
Db
         173 FLVNIATIFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPN--VFSM 230
QУ
                                                        : | |: : |||
                                ::||||||::::
                                             |: |
             11:
         171 FLYS-----PFDGILGLAFPSI---SAGGATTVMQKMLQENLLDFPVFSF 212
Db
         231 QMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNL 290
Qy
                        ]: || || || ||::|:|| | | :||: : |:|| |
         213 YLSGQ-----EGSQGGELVFGGVDPNLYTGQITWTPVTQTTYWQIGIEDFAVGGQSSGW 266
Db
         291 DCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSET 350
Qy
                   267 -CSQ--GCQGIVDTGTSLLTVPNQVFTELMQYIG----AQADD---SGQYVASCSNIE- 314
Db
         351 PWSYFPKI-----SIYLRDENS---SRSFRITILPQLYIQPMMGAGLNYECY 394
Qv
                                                  1 11
                                                          | |: || :
         315 ---YMPTITFVISGTSFPLPPSAYMLOSNSDYCTVGIESTYLPSQTGQPLW----- 362
Db
         395 RFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qy
                       ::| : :| |:|
                                          :1|||:
         363 -----ILGDVFLRVYYSIYDMGNNQVGFATA 388
Db
RESULT 6
A29937
gastricsin (EC 3.4.23.3) precursor - human
N; Alternate names: pepsin C; pepsinogen C
C; Species: Homo sapiens (man)
C;Date: 17-Oct-1988 #sequence revision 17-Oct-1988 #text change 31-Mar-2000
C; Accession: A29937; A31811; PX0028; I54213; A91125; A23458
R;Hayano, T.; Sogawa, K.; Ichihara, Y.; Fujii-Kuriyama, Y.; Takahashi, K.
J. Biol. Chem. 263, 1382-1385, 1988
A; Title: Primary structure of human pepsinogen C gene.
A; Reference number: A29937; MUID: 88087276; PMID: 3335549
A; Accession: A29937
A; Molecule type: DNA
A; Residues: 1-388 <HAY>
R; Taggart, R.T.; Cass, L.G.; Mohandas, T.K.; Derby, P.; Barr, P.J.; Pals, G.;
Bell, G.I.
J. Biol. Chem. 264, 375-379, 1989
A; Title: Human pepsinogen C (progastricsin). Isolation of cDNA clones,
localization to chromosome 6, and sequence homology with pepsinogen A.
A; Reference number: A31811; MUID: 89079679; PMID: 2909526
A; Accession: A31811
A; Molecule type: mRNA
A; Residues: 1-388 <TAG>
A;Cross-references: GB:J04443; NID:g551175; PIDN:AAA60074.1; PID:g551176
R; Athauda, S.B.P.; Tanji, M.; Kageyama, T.; Takahashi, K.
J. Biochem. 106, 920-927, 1989
A; Title: A comparative study on the NH2-terminal amino acid sequences and some
other properties of six isozymic forms of human pepsinogens and pepsins.
A; Reference number: PX0023; MUID: 90130402; PMID: 2515193
A; Accession: PX0028
A; Molecule type: protein
A; Residues: 17-101 <ATH>
```

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R; Pals, G.; Azuma, T.; Mohandas, T.K.; Bell, G.I.; Bacon, J.; Samloff, I.M.;
Walz, D.A.; Barr, P.J.; Taggart, R.T.
Genomics 4, 137-148, 1989
A; Title: Human pepsinogen C (progastricsin) polymorphism: evidence for a single
locus located at 6p21.1-pter.
A; Reference number: I54213; MUID: 89290840; PMID: 2567697
A; Accession: I54213
A; Status: translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-388 < RES>
A; Cross-references: GB:M23077; NID:g189830; PIDN:AAA60063.1; PID:g387015;
GB:J03063
A; Note: parts of this sequence, including the amino end and carboxyl ends of the
mature protein, were determined by protein sequencing
R; Foltmann, B.; Jensen, A.L.
Eur. J. Biochem. 128, 63-70, 1982
A; Title: Human progastricsin. Analysis of intermediates during activation into
gastricsin and determination of the amino acid sequence of the propart.
A; Reference number: A91125; MUID: 83079318; PMID: 6816595
A; Accession: A91125
A; Molecule type: protein
A; Residues: 17-39, 'ED', 42-51, 'S', 53-64 <FOL>
A; Note: pro-form; 29-Leu was also found
A; Note: activation at pH 2 is proposed to involve conformation change, cleavage
after Phe-42, and cleavage after Leu-59
C; Genetics:
A; Gene: GDB: PGC
A; Cross-references: GDB:119485; OMIM:169740
A; Map position: 6p21.3-6p21.1
A;Introns: 20/2; 70/3; 110/1; 149/3; 216/2; 256/2; 305/3; 338/3
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; hydrolase; protein digestion; stomach; zymogen
F;1-16/Domain: signal sequence #status predicted <SIG>
F;17-59/Domain: propeptide #status experimental <PRO>
F;60-388/Product: gastricsin #status experimental <MAT>
                         13.1%; Score 353; DB 2; Length 388;
  Query Match
  Best Local Similarity 29.1%; Pred. No. 1.1e-18;
  Matches 120; Conservative 65; Mismatches 120; Indels 108; Gaps
                                                                          21;
           52 PAERHADG-LALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVD 110
Qy
              1: 1: 111111 :1 1
           50 PAWKYRFGDLSVTYEP-----MAYMD-----AAYFGEISIGTPPQNFLVLFD 91
Db
          111 TGSSNFAV-----AGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDL 162
Qу
                            :::1 11 111 1 1
              11111
           92 TGSSNLWVPSVYCQSQACTSHS----RFNPSESSTYSTNGQTFSLQYGSGSLTGFFGYDT 147
Db
          163 VTIPKGFNTSFLVNIATIFESENFFLPG----IKWNGILGLAYATLAKPSSSLETFFDS 217
Qу
                                             :::||:|||| |:
                                                              :: |
                             1 1
          148 LTV----QSIQVPNQEFGLSEN--EPGTNFVYAQFDGIMGLAYPALSVDEAT--TAMQG 198
Db
          218 LVTQANIPN-VFSMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQI 276
Qу
                                      ]::||::| ||:: ||| | |:: |: ||
              :|:::||:::
          199 MVQEGALTSPVFSVYLSNQ-----QGSSGGAVVFGGVDSSLYTGQIYWAPVTQELYWQI 252
Db
          277 EILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGF 336
Qу
```

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253 GIEEFLIGGQASGW-CSE--GCQAIVDTGTSLLTVPQQYMSALLQA----- 295
Db
         337 WTGSOLACWTNSETPWSYF-----PKISIYLRDENSSRSFRITILPQLYIQPMMG 386
Qy
                      | : |
                                     | :: :
                                                       : | ||
         296 -TGAQ------EDEYGQFLVNCNSIQNLPSLTFII-----NGVEFPLPPSSYI---- 336
Db
         387 AGLNYECY-RFGISP----STNA---LVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qy
                     1: | | |
                                   ::| : :| ::|
         337 --LSNNGYCTVGVEPTYLSSQNGQPLWILGDVFLRSYYSVYDLGNNRVGFATA 387
Db
RESULT 7
JC7246
pepsinogen C - common marmoset
C; Species: Callithrix jacchus (common marmoset)
C;Date: 09-Jun-2000 #sequence revision 09-Jun-2000 #text change 21-Jul-2000
C:Accession: JC7246
R; Kageyama, T.
J. Biochem. 127, 761-770, 2000
A; Title: New world monkey pepsinogens A and C, and prochymosins. Purification,
characterization of enzymatic properties, cDNA cloning, and molecular evolution.
A; Reference number: JC7245
A; Accession: JC7246
A; Molecule type: mRNA
A; Residues: 1-388 < KAG>
A:Cross-references: DDBJ:AB038385
A; Experimental source: strain NW791
C; Comment: This protein, a zymogen of pepsins, is the major proteolytic enzyme
in vertebrate gastric juices. It plays roles in gastric digestion, and is a
useful molecular marker for clarifying the evolution of mammalian orders and
families.
C; Superfamily: pepsin
C; Keywords: gastric juice; zymogen
                       13.1%; Score 351.5; DB 2; Length 388;
 Query Match
 Best Local Similarity 30.1%; Pred. No. 1.4e-18;
 Matches 112; Conservative 56; Mismatches 115; Indels
                                                                    17;
          92 YYLEMLIGTPPQKLQILVDTGSSNFAV-----AGTPHSYIDTYFDTERSSTYRSKGF 143
Qу
             |: |: ||||| :| |||||| |
                                            1 1 11
                                                     |:
          73 YFGEISIGTPPQNFLVLFDTGSSNLWVPSVYCQSQACTSHS----RFNPSASSTYSSNGQ 128
Db
         144 DVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFESENFFLPG----IKWNGILG 198
Qу
              :::| || || || || || ||
                                               129 TFSLQYGSGSLTGFFGYDTLTV-----QSIQVPNQEFGLSEN---EPGTNFVYAQFDGIMG 181
Db
         199 LAYATLAKPSSSLETFFDSLVTQANIPN-VFSMQMCGAGLPVAGSGTNGGSLVLGGIEPS 257
Qу
                    182 LAYPALSMGGAT--TAMQGMLQEGALTSPVFSFYLSNQ-----QGSSGGAVIFGGVDSS 233
Db
         258 LYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFD 317
Qу
             234 LYTGQIYWAPVTQELYWQIGIEEFLIGGQASGW-CSE--GCQAIVDTGTSLLTVPQQYMS 290
Db
         318 AVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYF-----PKISIYLRDENS 367
Qу
                               11:1 : 1
             |:11
```

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291 AFLEA----TGAQ-----EDEYGQFLVNCDSIQNLPTLTFII---- 323
Db
         368 SRSFRITILPQLYIQPMMGAGLNYECY-RFGISP-----STNALVIGATVMEGFYVIF 419
Qy
                                                      ::| : :| :|
                   : | ||
                                1: | |: |
                                                  - 1
         324 -NGVEFPLPPSSYI-----LSNNGYCTVGVEPTYLSSQNSQPLWILGDVFLRSYYSVF 375
Db
         420 DRAQKRVGFAAS 431
Qу
                 11111:
             1
         376 DLGNNRVGFATA 387
Db
RESULT 8
B43356
gastricsin (EC 3.4.23.3) precursor - guinea pig
N; Alternate names: pepsin C
C; Species: Cavia porcellus (guinea pig)
C;Date: 03-Feb-1994 #sequence revision 03-Feb-1994 #text change 22-Jun-1999
C:Accession: B43356
R; Kageyama, T.; Ichinose, M.; Tsukada, S.; Miki, K.; Kurokawa, K.; Koiwai, O.;
Tanji, M.; Yakabe, E.; Athauda, S.B.; Takahashi, K.
J. Biol. Chem. 267, 16450-16459, 1992
A; Title: Gastric procathepsin E and progastricsin from guinea pig. Purification,
molecular cloning of cDNAs, and characterization of enzymatic properties, with
special reference to procathepsin E.
A; Reference number: A43356; MUID: 92355614; PMID: 1644829
A; Accession: B43356
A; Status: preliminary
A; Molecule type: mRNA
A; Residues: 1-394 < KAG>
A;Cross-references: GB:M88652; NID:g191296; PIDN:AAA37053.1; PID:g191297
A; Note: sequence extracted from NCBI backbone (NCBIN:110805, NCBIP:110806)
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; gastric juice; hydrolase; protein digestion;
stomach
                        12.1%; Score 324.5; DB 2; Length 394;
  Query Match
  Best Local Similarity 29.0%; Pred. No. 1.5e-16;
  Matches 107; Conservative 63; Mismatches 116; Indels
                                                            83; Gaps
                                                                       18;
          92 YYLEMLIGTPPQKLQILVDTGSSNF-----AVAGTPHSYIDTYFDTERSSTYRSKGF 143
Qу
             ::| | |
                                                     1 1:
          79 YFGQISLGTPPQSFQVLFDTGSSNLWVPSVYCSSLACTTH----TRFNPRDSSTYVATDQ 134
Db
         144 DVTVKYTOGSWTGFVGEDLVTI----PK-GFNTSFLVNIATIFESENFFLPG----IK 192
Qу
               :::| || || || || || ||
                                     |:| ||
         135 SFSLEYGTGSLTGVFGYDTMTIQDIQVPKQEFGLS-----ETE----PGSDFVYAE 181
Db
         193 WNGILGLAYATLAKPSSSLETFFDSLVTQANI-PNVFSMQMCGAGLPVAGS--GTNGGSL 249
Qу
             ::||||| | |:: :: | |:: ::||:: | || |:: | |
         182 FDGILGLGYPGLSEGGAT--TAMQGLLREGALSQSLFSVYL-----GSQQGSDEGQL 231
Db
         250 VLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLL 309
Qу
             232 ILGGVDESLYTGDIYWTPVTQELYWQIGIEGFLIDGSASGWCSR---GCQGIVDTGTSLL 288
Db
         310 RLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSR 369
Qу
                     :1:1:
                                1:: 1
                                          ::|::|:::
              : |
```

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289 TVPSDYLSTLVQAIGAEE--NEYGEYF-----VSCSSIQDLPTLTFVISGV------ 332
Db
         370 SFRITILPQLYIQP----MMGAGLNYECYRFGISPSTN--ALVIGATVMEGFYVIFDRA 422
Qy
                  : 1 11
                               1:1
                                     : 11
                                                      ::|
                                                           : :| ::| |
         333 -- EFPLSPSAYILSGENYCMVGLESTY-----VSPGGGEPVWILGDVFLRSYYSVYDLA 384
Db
         423 QKRVGFAAS 431
Qу
               11111:
         385 NNRVGFATA 393
Db
RESULT 9
JC7575
pepsinogen A - bullfrog
C; Species: Rana catesbeiana (bullfrog)
C;Date: 30-Jun-2001 #sequence revision 30-Jun-2001 #text change 03-Aug-2001
C:Accession: JC7575
R; Ikuzawa, M.; Inokuchi, T.; Kobayashi, K.; Yasumasu, S.
J. Biochem. 129, 147-153, 2001
A; Title: Amphibian pepsinogens: Purification and characterization of Xenopus
pepsinogens, and molecular cloning of Xenopus and bullfrog pepsinogens.
A; Reference number: JC7573; MUID:21064922; PMID:11134969
A; Contents: Stomach
A; Accession: JC7575
A; Molecule type: mRNA
A; Residues: 1-385 < IKU>
A; Cross-references: DDBJ:AB045376
C: Comment: This protein is a zymogen for gastric aspartic proteinase, with
pepsin-like activity.
C; Genetics:
A; Gene: PqA
C; Superfamily: pepsin
C; Keywords: stomach; zymogen
                        11.9%; Score 320; DB 2; Length 385;
  Query Match
  Best Local Similarity 27.8%; Pred. No. 3.2e-16;
  Matches 111; Conservative 67; Mismatches 147; Indels
                                                            74; Gaps
                                                                         15;
          50 GTPAERHADGLALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILV 109
Qу
                     1 ::1
          39 GDYLKKHHYNPATKYFPSLAQASG-----EPLQNYMDIEYFGTISIGTPPQSFTVIF 90
Db
         110 DTGSSNFAVAGTPHSYIDT-----YFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDL 162
Qy
                                       1: ::|||:::
                                                      1:::| || :||:| |
             91 DTGSSNLWV---PSVYCSSPACTNHHMFNPQQSSTFQATNTPVSIQYGTGSMSGFLGYDT 147
Db
         163 VTIPKGFNTSFLVNIATIFESE-NFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQ 221
Qу
                          | | : :|| || ::|||||: :|| ||
                                                                11:: 1
         148 VOVG---NIOITNOIFGLSOSEPGSFLYYSPFDGILGLAFPSLA--SSQATPVFDNMWNQ 202
Db
         222 ANIP-NVFSMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYOIEILK 280
Qy
                              : | :| :: ||:: | | |:: : |: | |:|| :
         203 GLIPODLFSVYL-----SSQGQSGSFVLFGGVDTSYYTGNLNWVPLTAETYWQITVDS 255
Db
         281 LEIGGOSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGS 340
Qγ
                            : ||||:||:||
                                               | :: |:
         256 ISIGGQVIACS----GSCSAIVDTGTSLLAGPSTPI-ANIQYYIGAN---QDSNGQYV-- 305
Db
```

```
341 OLACWTNSETPWSYFP-----KISIYLRDENSS--RSFRITILPQLYIQPMMGAGLN 390
Qу
                   | |:|
                                                ı
                                                      1:
                                                          - 11
          306 -INCNNISNMPTVVFTINGVQYPLPASAYVRQSQQSCTSGFQAMNLP----- 351
Db
          391 YECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFA 429
Qy
                      : | : ::| : :||:|||
                                                1 1
          352 -----TSSGDLWILGDVFIREYYVVFDRANNYVAMA 382
Dh
RESULT 10
REMSK
renin (EC 3.4.23.15) precursor, renal - mouse
N; Alternate names: angiotensin-forming enzyme; angiotensinogenase; renin 1
C; Species: Mus musculus (house mouse)
C; Date: 30-Jun-1987 #sequence revision 30-Jun-1987 #text change 18-Jun-1999
C; Accession: A00989; S07636; A22766; A22058; I57576; A05137; JH0083
R; Holm, I.; Ollo, R.; Panthier, J.J.; Rougeon, F.
EMBO J. 3, 557-562, 1984
A; Title: Evolution of aspartyl proteases by gene duplication: the mouse renin
gene is organized in two homologous clusters of four exons.
A; Reference number: A00989; MUID: 84182525; PMID: 6370686
A; Accession: A00989
A; Molecule type: DNA
A; Residues: 1-402 <HOL>
A; Cross-references: EMBL:X00850
R; Kim, W.S.; Murakami, K.; Nakayama, K.
Nucleic Acids Res. 17, 9480, 1989
A; Title: Nucleotide sequence of a cDNA coding for mouse Ren1 preprorenin.
A; Reference number: S07636; MUID: 90067953; PMID: 2685761
A; Accession: S07636
A; Molecule type: mRNA
A; Residues: 1-402 <KIM>
A; Cross-references: EMBL: X16642; NID: g53930; PIDN: CAA34636.1; PID: g53931
R; Mullins, J.J.; Burt, D.W.; Windass, J.D.; McTurk, P.; George, H.; Brammar,
W.J.
EMBO J. 1, 1461-1466, 1982
A; Title: Molecular cloning of two distinct renin genes from the DBA/2 mouse.
A; Reference number: A90968; MUID: 84207899; PMID: 6327270
A; Accession: A22766
A; Molecule type: mRNA
A; Residues: 269-314, 'D', 316 < MUL>
R; Panthier, J.J.; Dreyfus, M.; Roux, D.T.L.; Rougeon, F.
Proc. Natl. Acad. Sci. U.S.A. 81, 5489-5493, 1984
A; Title: Mouse kidney and submaxillary gland renin genes differ in their 5'
putative regulatory sequences.
A; Reference number: A22058; MUID: 84298161; PMID: 6089205
A: Accession: A22058
A; Molecule type: DNA
A; Residues: 1-30 < PAN>
R; Field, L.J.; Philbrick, W.M.; Howles, P.N.; Dickinson, D.P.; McGowan, R.A.;
Gross, K.W.
Mol. Cell. Biol. 4, 2321-2331, 1984
A; Title: Expression of tissue-specific Ren-1 and Ren-2 genes of mice:
Comparative analysis of 5'-proximal flanking regions.
A; Reference number: 157576; MUID: 85085936; PMID: 6392850
A; Accession: I57576
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A; Status: preliminary; translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-31 < RES>
A;Cross-references: GB:K02800; NID:q200689; PIDN:AAA40044.1; PID:q200690
C; Comment: The only known function of renal renin is to release angiotensin I
from angiotensinogen in the plasma, initiating a cascade of reactions that
produces an elevation of blood pressure and increased sodium retention by the
kidney.
C; Comment: Renal renin is synthesized by the juxtaglomerular cells of the kidney
in response to decreased blood pressure and sodium concentration.
C; Genetics:
A; Gene: Ren-1
A; Introns: 31/2; 81/3; 123/1; 162/3; 228/2; 268/2; 316/3; 349/3
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; blood pressure control; glycoprotein;
hydrolase; kidney; plasma
F;1-21/Domain: signal sequence #status predicted <SIG>
F:22-64/Domain: propeptide #status predicted <PRO>
F:65-402/Product: renin #status predicted <MAT>
F;69,139,320/Binding site: carbohydrate (Asn) (covalent) #status predicted
F;102,287/Active site: Asp #status predicted
                        11.9%; Score 320; DB 1; Length 402;
  Query Match
  Best Local Similarity 28.6%; Pred. No. 3.4e-16;
 Matches 126; Conservative 66; Mismatches 181; Indels 68; Gaps
                                                                       21;
          10 LPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPG-PGTPAERHADGLALALE--- 65
Qу
                          :| |:|| |
             :11 | 11
                                           : |
           6 MPLWALLLL----WSPCTFSLPTRTATFERIPLKKMPSVREILEERGVDMTRLSAEWGV 60
Db
          66 ----PA---LASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAV 118
Qу
                                  1.1
                                           1:
                     1 11
                            1:1
          61 FTKRPSLTNLTSPVVLTNYL----NTQ------YYGEIGIGTPPQTFKVIFDTGSANLWV 110
Db
         119 AGTPHSY-----IDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTS 172
Qv
               | |
                        | : ::: ||:| | | | | |
                                       | | | |: | |
                                                    | | | : : | | | | : | |
         111 PSTKCSRLYLACGIHSLYESSDSSSYMENGSDFTIHYGSGRVKGFLSQDSVTV-GGITVT 169
Db
         173 FLVNIATIFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANI-PNVFSMQ 231
Qy
                  170 QTFGEVTELPLIPFML--AKFDGVLGMGFP--AQAVGGVTPVFDHILSQGVLKEEVFSVY 225
Db
         232 MCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLD 291
Qy
                          \Pi
         226 Y----NRGSHLLGGEVVLGGSDPOHYQGNFHYVSISKTDSWQITMKGVSVG--SSTLL 277
Db
         292 CREYNADKAIVDSGTTLLRLPQKVFDAVVEAV-ARASLIPEFSDGFWTGSQLACWTNSET 350
Qy
             | | | :||:|:: | :::|: |: | |::
                                                            : |
         278 CEEGCA--VVVDTGSSFISAPTSSLKLIMQALGAKEKRIEEY-----VVNC---SQV 324
Db
         351 PWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGL-NYECYRFGISPSTNAL-VIG 408
QУ
                                      : 1
                                               1
                              1:: ::
         325 P--TLPDISFDL----GGRAYTLSSTDYVLQYPNRRDKLCTLALHAMDIPPPTGPVWVLG 378
Db
         409 ATVMEGFYVIFDRAQKRVGFA 429
Qу
             11: 11 111
                            1:11
         379 ATFIRKFYTEFDRHNNRIGFA 399
Db
```

```
RESULT 11
S66516
oryzasin (EC 3.4.23.-) precursor - rice
N; Alternate names: aspartic proteinase 1
C; Species: Oryza sativa (rice)
C;Date: 28-Oct-1996 #sequence revision 13-Mar-1997 #text change 20-Jun-2000
C; Accession: S66516; S66517
R; Asakura, T.; Watanabe, H.; Abe, K.; Arai, S.
Eur. J. Biochem. 232, 77-83, 1995
A; Title: Rice aspartic proteinase, oryzasin, expressed during seed ripening and
germination, has a gene organization distinct from those of animal and microbial
aspartic proteinases.
A; Reference number: S66516; MUID: 96048031; PMID: 7556174
A:Accession: S66516
A; Molecule type: DNA
A; Residues: 1-509 < ASA>
A;Cross-references: EMBL:D32165; NID:g511665; PIDN:BAA06876.1; PID:q1030715
A; Accession: S66517
A; Molecule type: mRNA
A; Residues: 1-509 <ASZ>
A;Cross-references: EMBL:D32144; NID:g1255684; PIDN:BAA06875.1; PID:g1711289
C; Comment: The pair of saposin repeat homology domains tagged SAP1 and SAP2
represent a cyclical permutation of a single saposin repeat.
C:Genetics:
A; Introns: 119/3; 140/1; 171/3; 209/2; 265/3; 279/1; 300/3; 338/3; 360/2; 412/3;
452/3; 482/2
C; Superfamily: oryzasin; saposin repeat homology
C; Keywords: aspartic proteinase; hydrolase
F;1-20/Domain: signal sequence #status predicted <SIG>
F;21-68/Domain: propeptide #status predicted <PRO>
F;68-509/Product: aspartic proteinase 1 #status predicted <MAT>
F;316-361/Domain: saposin repeat homology #status atypical <SAP1>
F;370-420/Domain: saposin repeat homology #status atypical <SAP2>
F;103,290/Active site: Asp #status predicted
                         11.7%; Score 313.5; DB 2; Length 509;
  Query Match
  Best Local Similarity 23.0%; Pred. No. 1.5e-15;
 Matches 127; Conservative 75; Mismatches 179; Indels 171; Gaps
                                                                         19;
            3 ALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGLAL 62
Qу
                                                            :11 | |
             ::| || :| | || :|
                                        5 SVALVLLAAVLLQALLPASAEEGLVRIALKKRPIDENSRVAARLSG----EEGARRLGL 59
Db
           63 ALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSN----- 115
Qу
                       60 RGANSLGGGGGGGDIVALKNYMNAQ----YFGEIGVGTPPQKFTVIFDTGSSNLWVPSAK 115
Db
          116 --FAVAGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSF 173
Qу
                                            ::| || || || || ||
                            :::||||: |
          116 CYFSIACFFHS----RYKSGQSSTYQKNGKPAAIQYGTGSIAGFFSEDSVTVGD----- 165
Dh
          174 LVNIATIFESENFF----LPGI-----KWNGILGLAYATLAKPSSSLETFFDSLVTQANI 224
Qу
                                        1::||||| : :: :
                   ::: | | | | | | |
          166 -----LVVKDQEFIEATKEPGLTFMVAKFDGILGLGFQEISVGDA------V 206
Db
```

```
225 PNVFSMQMCG-AGLPVAGSGTN-----GGSLVLGGIEPSLYKGDIWYTPIKEEWYYQI 276
Qy
            207 PVWYKMVEQGLVSEPVFSFWFNRHSDEGEGGEIVFGGMDPSHYKGNHTYVPVSQKGYWQF 266
Db
         277 EILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPE---- 331
Qу
            267 EMGDVLIGGKTTGF-CA--SGCSAIADSGTSLLAGPTAIITEINEKIGATGVVSQECKTV 323
Db
         332 -----FSDGF----- 336
Qу
                                                         1:1
         324 VSQYGQQILDLLLAETQPSKICSQVGLCTFDGKHGVSAGIKSVVDDEAGESNGLQSGPMC 383
Db
         337 -----FPKISIYLRD 364
Qу
                   1:111 : :1
                                                         |:|| :
         384 NACEMAVVWMQNQLAQNKTQDLILNYINQLCDKLPSPMGESSVDCGSLASMPEISFTIGA 443
Db
         365 ENSSRSFRITILPQLYIQPMMGAGLNYECY----RFGISPSTNAL-VIGATVMEGFYVIF 419
Qу
            444 K-----KFALKPEEYIL-KVGEGAAAQCISGFTAMDIPPPRGPLWILGDVFMGAYHTVF 496
Db
Qу
         420 DRAQKRVGFAAS 431
            \perp: | | | | | |
Db
         497 DYGKMRVGFAKS 508
RESULT 12
A24608
qastricsin (EC 3.4.23.3) precursor - rat
N; Alternate names: pepsinogen C
N; Contains: pepsin A (EC 3.4.23.1) precursor
C; Species: Rattus norvegicus (Norway rat)
C;Date: 30-Jun-1988 #sequence revision 05-Aug-1994 #text change 18-Jun-1999
C; Accession: A33510; A24608; C22434; A05145; A61298
R; Ishihara, T.; Ichihara, Y.; Hayano, T.; Katsura, I.; Sogawa, K.; Fujii-
Kuriyama, Y.; Takahashi, K.
J. Biol. Chem. 264, 10193-10199, 1989
A; Title: Primary structure and transcriptional regulation of rat pepsinogen C
gene.
A; Reference number: A33510; MUID: 89255508; PMID: 2722863
A; Accession: A33510
A; Molecule type: DNA
A; Residues: 1-392 <ISH>
A; Cross-references: GB:M25985
R;Ichihara, Y.; Sogawa, K.; Morohashi, K.; Fujii-Kuriyama, Y.; Takahashi, K.
Eur. J. Biochem. 161, 7-12, 1986
A; Title: Nucleotide sequence of a nearly full-length cDNA coding for pepsinogen
of rat gastric mucosa.
A; Reference number: A24608; MUID: 87054020; PMID: 3780741
A; Accession: A24608
A; Molecule type: mRNA
A; Residues: 1-392 <ICH>
A;Cross-references: GB:X04644; NID:g56880; PIDN:CAA28305.1; PID:g56881
R; Ichihara, Y.; Sogawa, K.; Takahashi, K.
J. Biochem. 98, 483-492, 1985
A; Title: Isolation of human, swine, and rat prepepsinogens and calf
preprochymosin, and determination of the primary structures of their NH2-
terminal signal sequences.
```

```
A; Reference number: A22434; MUID: 86059312; PMID: 2415509
A; Accession: C22434
A; Molecule type: protein
A; Residues: 1-19, 'X', 21-23, 'X', 25-29 <IC2>
R; Arai, K.M.; Muto, N.; Tani, S.; Akahane, K.
Biochim. Biophys. Acta 788, 256-261, 1984
A; Title: The N-terminal sequence of rat pepsinogen.
A; Reference number: A05145; MUID: 84257697; PMID: 6743670
A; Accession: A05145
A; Molecule type: protein
A; Residues: 17-30, 'Q', 32-102, 'A', 104-108, 'L', 110-112 <ARA>
A; Experimental source: Wistar strain
R; Ichihara, Y.; Sogawa, K.; Takahashi, K.
J. Biochem. 92, 603-606, 1982
A; Title: Rat gastric prepepsinogen: in vitro synthesis and partial amino-
terminal signal sequence.
A; Reference number: A61298; MUID: 83030750; PMID: 6182139
A:Accession: A61298
A; Molecule type: protein
A; Residues: 1, 'XX', 4-6, 'X', 8-9, 'X', 11, 'X', 13-14, 'XXX', 18-
19,'X',21,'X',23,'XX',26,'X' <IC3>
C; Comment: This enzyme has more restricted specificity than pepsin A. It is the
major form of pepsinogen in rat gastric mucosa.
C; Genetics:
A; Introns: 20/2; 73/3; 113/1; 152/3; 219/2; 259/2; 309/3; 342/3
A; Note: there are at least two very similar genes for gastricsin in rat
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; gastric juice; hydrolase; protein digestion;
stomach
F;1-16/Domain: signal sequence #status experimental <SIG>
F:17-392/Product: pepsinogen #status experimental <MAT>
F;17-62/Domain: activation peptide #status experimental <ACT>
F;94,280/Active site: Asp #status predicted
F;107-112,270-275,314-347/Disulfide bonds: #status predicted
                          11.6%; Score 313; DB 1; Length 392;
  Query Match
  Best Local Similarity 29.5%; Pred. No. 1.1e-15;
  Matches 105; Conservative 56; Mismatches 139; Indels
                                                                           16;
           92 YYLEMLIGTPPQKLQILVDTGSSNFAV-----AGTPHSYIDTYFDTERSSTYRSKGF 143
Qу
              1: 1: |||||| :| ||||||
                                                1 | 1:
                                                          |: :|||| ::|
           76 YFGEISIGTPPQNFLVLFDTGSSNLWVSSVYCQSEACTTHA----RFNPSKSSTYYTEGQ 131
Db
          144 DVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFESENFFLPG----IKWNGILG 198
Qу
                                                         11
                1 1
                                                    132 TFSLQYGTGSLTGFFGYDTLTV----QSIQVPNQEFGLSEN--EPGTNFVYAQFDGIMG 184
Db
          199 LAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVAGS--GTNGGSLVLGGIEP 256
Qу
                                          ::::
                  1: 1
                           1
          185 LAYPGLS--SGGATTALQGMLGE----GALSQPLFGVYL---GSQQGSNGGQIVFGGVDK 235
Db
          257 SLYKGDIWYTPIKEEWYYOIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVF 316
Qγ
              : | | | | : | : | | | | | | |
                                       11 1:
                                               1
                                                      : |||:||:|| :| :
          236 NLYTGEITWVPVTQELYWQITIDDFLIGDQASGW-CSSQGC-QGIVDTGTSLLVMPAQYL 293
Db
          317 DAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITIL 376
Qу
                            |::|
                                      ::|:|
                                                    1:1
                ::: :
```

```
294 SELLQTIGAQE--GEYGEYF-----VSCDSVSS-----LPTLSFVL-----NGVQFPLS 335
Db
         377 POLY-IOPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qy
                    : 1
                                       ::|
                                           : :| |||
            1 1 1
Db
         336 PSSYIIQEDNFCMVGLESISLTSESGQPLWILGDVFLRSYYAIFDMGNNKVGLATS 391
RESULT 13
A41443
pepsin (EC 3.4.23.-) precursor, embryonic - chicken
C; Species: Gallus gallus (chicken)
C;Date: 05-Jun-1992 #sequence revision 05-Jun-1992 #text change 21-Jul-2000
C; Accession: A41443
R; Hayashi, K.; Aqata, K.; Mochii, M.; Yasuqi, S.; Equchi, G.; Mizuno, T.
J. Biochem. 103, 290-296, 1988
A; Title: Molecular cloning and the nucleotide sequence of cDNA for embryonic
chicken pepsinogen: phylogenetic relationship with prochymosin.
A: Reference number: A41443; MUID: 88227903; PMID: 3131317
A:Accession: A41443
A; Status: preliminary
A; Molecule type: mRNA
A; Residues: 1-383 <HAY>
A;Cross-references: GB:D00215; NID:g2760810; PIDN:BAA00153.1; PID:g222853
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; hydrolase; protein digestion
 Query Match
                       11.5%; Score 310; DB 2; Length 383;
 Best Local Similarity 26.8%; Pred. No. 1.8e-15;
 Matches 106; Conservative 63; Mismatches 136; Indels
                                                                    15:
         56 HA--DGLALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGS 113
Qy
               11: 111
                             1 1 1
                                             Db
         55 HAFPDVLTVVTEPLL-----NTLDM-----EYYGTISIGTPPQDFTVVFDTGS 97
         114 SNFAVAG----TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGF 169
QУ
            || :|
                             98 SNLWVPSVSCTSPACQSHQMFNPSQSSTYKSTGQNLSIHYGTGDMEGTVGCDTVTVASLM 157
Db
         170 NTSFLVNIATIFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANI-PNVF 228
Qу
            158 DTNQLFGLST-SEPGQFFV-YVKFDGILGLGYPSLA--ADGITPVFDNMVNESLLEQNLF 213
Db
         229 SMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSL 288
Qy
                          _ | :| |||: | : | | : |: |:|| : : : |:
            1:::
         214 SVYLSREPM----GSMVVFGGIDESYFTGSINWIPVSYQGYWQISMDSIIVNKQEI 265
Db
         289 NLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNS 348
Qy
                  : :||:|:|:|: | : : ||
         266 ACS----SGCQAIIDTGTSLVAGPASDINDIQSAVG------ANQ 300
Db
         349 ETPWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGLNY-----ECY 394
Qy
                           301 NTYGEY----SVNCSHILAMPDVVF--VIG-GIOYPVPALAYTEQNGQGTCM 345
Db
         395 RFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFA 429
Qy
                346 SSFQNSSADLWILGDVFIRVYYSIFDRANNRVGLA 380
Dh
```

```
RESULT 14
KHHUD
cathepsin D (EC 3.4.23.5) precursor [validated] - human
N; Alternate names: preprocathepsin D
C; Species: Homo sapiens (man)
C;Date: 28-Dec-1987 #sequence revision 28-Dec-1987 #text_change 15-Sep-2000
C; Accession: A25771; S30749; PC2066; I59236; I57716
R; Faust, P.L.; Kornfeld, S.; Chirgwin, J.M.
Proc. Natl. Acad. Sci. U.S.A. 82, 4910-4914, 1985
A; Title: Cloning and sequence analysis of cDNA for human cathepsin D.
A; Reference number: A25771; MUID: 85270436; PMID: 3927292
A; Accession: A25771
A; Molecule type: mRNA
A; Residues: 1-412 <FAU>
A; Cross-references: EMBL: M11233; NID: q181179; PIDN: AAB59529.1; PID: q181180
R; Westley, B.R.; May, F.E.B.
Nucleic Acids Res. 15, 3773-3786, 1987
A:Title: Oestrogen regulates cathepsin D mRNA levels in oestrogen responsive
human breast cancer cells.
A; Reference number: S30749; MUID: 87231068; PMID: 3588310
A; Accession: S30749
A; Molecule type: mRNA
A; Residues: 1-412 <WES>
A; Cross-references: EMBL: X05344; NID: g29677; PIDN: CAA28955.1; PID: g29678
R; May, F.E.B.; Smith, D.J.; Westley, B.R.
Gene 134, 277-282, 1993
A; Title: The human cathepsin D-encoding gene is transcribed from an estrogen-
regulated and a constitutive start point.
A; Reference number: PC2066; MUID: 94085791; PMID: 8262386
A; Accession: PC2066
A; Molecule type: DNA
A; Residues: 1-23 <MAY>
A;Cross-references: GB:L12980; NID:q291930; PIDN:AAA16314.1; PID:q455429
A; Experimental source: MCF-7 cell
R; Cavailles, V.; Augereau, P.; Rochefort, H.
Proc. Natl. Acad. Sci. U.S.A. 90, 203-207, 1993
A; Title: Cathepsin D gene is controlled by a mixed promoter, and estrogens
stimulate only TATA-dependent transcription in breast cancer cells.
A; Reference number: 159236; MUID: 93126342; PMID: 8419924
A; Accession: I59236
A; Status: translation not shown; translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-22 <CAV1>
A;Cross-references: GB:S52557; NID:q263124; PIDN:AAD13868.1; PID:q4261568
R; Augereau, P.; Miralles, F.; Cavailles, V.; Gaudelet, C.; Parker, M.;
Rochefort, H.
Mol. Endocrinol. 8, 693-703, 1994
A; Title: Characterization of the proximal estrogen-responsive element of human
cathepsin D gene.
A; Reference number: I57716; MUID: 95021301; PMID: 7935485
A; Accession: I57716
A; Status: translation not shown; translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-22 <CAV2>
A; Cross-references: GB: S74689; NID: q786350; PIDN: AAD14156.1; PID: q4261856
```

```
R; Baldwin, E.T.; Bhat, T.N.; Gulnik, S.; Erickson, J.W.
submitted to the Brookhaven Protein Data Bank, April 1993
A; Reference number: A51839; PDB:1LYA
A; Contents: annotation; X-ray crystallography, 2.5 angstroms, residues 65-
161;170-241
R; Baldwin, E.T.; Bhat, T.N.; Gulnik, S.; Erickson, J.W.
submitted to the Brookhaven Protein Data Bank, April 1993
A; Reference number: A51840; PDB:1LYB
A; Contents: annotation; X-ray crystallography, 2.5 angstroms, with inhibitor
residues 65-161;170-241
R; Baldwin, E.T.; Bhat, T.N.; Gulnik, S.; Hosur, M.V.; Sowder II, R.C.; Cachau,
R.E.; Collins, J.; Silva, A.M.; Erickson, J.W.
Proc. Natl. Acad. Sci. U.S.A. 90, 6796-6800, 1993
A; Title: Crystal structures of native and inhibited forms of human cathepsin D:
implications for lysosomal targeting and drug design.
A; Reference number: A48229; MUID: 93342076; PMID: 8393577
A; Contents: annotation; X-ray crystallography, 2.5 angstroms
C: Comment: Cathepsin D is a ubiquitous lysosomal proteinase.
C; Comment: In addition to the propeptide, residues 163-168 and 411-412 are
proteolytically removed. Residues 169 and 170 are also partially removed.
C; Comment: The carbohydrate bound to 134-Asn contains a mannose-6-phosphate that
is bound near 267-Lys and the phosphotransferase recognition region.
C; Genetics:
A:Gene: GDB:CTSD
A; Cross-references: GDB:120512; OMIM:116840
A; Map position: 11p15.5-11p15.5
C; Function:
A; Description: limited specificity endopeptidase
A; Pathway: intracellular protein degradation
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; glycoprotein; hydrolase; lysosome; protein
degradation
F;1-20/Domain: signal sequence #status predicted <SIG>
F;21-64/Domain: propeptide #status predicted <PRO>
F;65-162,169-410/Product: cathepsin D #status experimental <MAT>
F;267,329-356/Region: phosphotransferase recognition
F;91-160,110-117,286-290,329-366/Disulfide bonds: #status experimental
F;97,295/Active site: Asp #status experimental
F;134,263/Binding site: carbohydrate (Asn) (covalent) #status experimental
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  Matches 121; Conservative 75; Mismatches 180; Indels
                                                               71; Gaps
                                                                            22;
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Qу
                                          : | :: |
                       1 | | | | |
                                   : | |
            6 LLPLAL--CLLAAP--ASALVRIPLHKFTSIRRTMSEVGGSVEDLIAKGPVSKYSQAVPA 61
Db
           63 ALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGTP 122
Qу
                                           |::
                | ; |
           62 VTEGPI--PEVLKNYM------DAQYYGEIGIGTPPQCFTVVFDTGSSNLWVPSIH 109
Db
          123 HSYIDT-----YFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIP--KGFNTSFL 174
Qу
                                           : | || :|:: :| |::|
                          :::::||||
                                     1
          110 CKLLDIACWIHHKYNSDKSSTYVKNGTSFDIHYGSGSLSGYLSQDTVSVPCQSASSASAL 169
Db
          175 --VNIATIFESENFFLPGI----KWNGILGLAYATLAKPSSSLETFFDSLVTQANI-PN 226
Qу
```

```
1::||1|:|| :: :::
                                                            11:1: | : |
                            170 GGVKVEROVFGEATKQPGITFIAAKFDGILGMAYPRIS--VNNVLPVFDNLMQQKLVDQN 227
Db
          227 VFSMOMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQ 286
Qу
                                228 IFSFY----LSRDPDAQPGGELMLGGTDSKYYKGSLSYLNVTRKAYWQVHLDQVEV-AS 281
Db
          287 SLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWT 346
Qу
                         :|||:|:|:
                                          : |
          282 GLTL-CKE--GCEAIVDTGTSLMVGP---VDEVRELQKAIGAVPLIQGEY----MIPC-- 329
Db
          347 NSETPWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRF---GISPSTN 403
Qу
                                   ::::: |: | : ||
                   | | |:: |
                                                              - 1
          330 ---EKVSTLPAITLKL----GGKGYKLS--PEDYTLKVSQAGKTLCLSGFMGMDIPPPSG 380
Db
          404 AL-VIGATVMEGFYVIFDRAQKRVGFA 429
Qу
               11111
          381 PLWILGDVFIGRYYTVFDRDNNRVGFA 407
Db
RESULT 15
KHMSD
cathepsin D (EC 3.4.23.5) precursor - mouse
C; Species: Mus musculus (house mouse)
C;Date: 31-Dec-1991 #sequence revision 31-Dec-1991 #text change 18-Jun-1999
C; Accession: I48278; S14704; S12587
R; Hetman, M.; Perschl, A.; Saftig, P.; Von Figura, K.; Peters, C.
DNA Cell Biol. 13, 419-427, 1994
A; Title: Mouse cathepsin D gene: molecular organization, characterization of the
promoter, and chromosomal localization.
A; Reference number: 148278; MUID: 94280622; PMID: 8011168
A; Accession: I48278
A; Status: preliminary; translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-410 < RES>
A; Cross-references: EMBL: X68378; NID: q50302; PIDN: CAA48453.1; PID: q817945
R; Diedrich, J.F.; Staskus, K.A.; Retzel, E.F.; Haase, A.T.
Nucleic Acids Res. 18, 7184, 1990
A; Title: Nucleotide sequence of a cDNA encoding mouse cathepsin D.
A; Reference number: S14704; MUID: 91088345; PMID: 2263503
A; Accession: S14704
A; Molecule type: mRNA
A; Residues: 1-410 <DIE>
A; Cross-references: EMBL: X53337; NID: g50300; PIDN: CAA37423.1; PID: g50301
R; Grusby, M.J.; Mitchell, S.C.; Glimcher, L.H.
Nucleic Acids Res. 18, 4008, 1990
A; Title: Molecular cloning of mouse cathepsin D.
A; Reference number: S12587; MUID: 90326544; PMID: 2374732
A; Accession: S12587
A; Molecule type: mRNA
A; Residues: 1-410 <GRU>
A; Cross-references: EMBL: X52886; NID: g50298; PIDN: CAA37067.1; PID: g50299
C; Genetics:
A; Introns: 23/2; 76/3; 118/1; 157/3; 233/2; 274/2; 322/3; 355/3
C; Function:
A; Description: limited specificity endopeptidase
A; Pathway: intracellular protein degradation
```

```
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; glycoprotein; hydrolase; lysosome; protein
F;1-20/Domain: signal sequence #status predicted <SIG>
F;21-64/Domain: propeptide #status predicted <PRO>
F;65-410/Product: cathepsin D, single-chain form #status predicted <MAT>
F;91-160,110-117,284-288,327-364/Disulfide bonds: #status predicted
F;97,293/Active site: Asp #status predicted
F;134,261/Binding site: carbohydrate (Asn) (covalent) #status predicted
                      11.4%; Score 306.5; DB 1; Length 410;
 Query Match
 Best Local Similarity 27.5%; Pred. No. 3.6e-15;
 Matches 103; Conservative 64; Mismatches 123; Indels 85; Gaps
                                                                15;
         92 YYLEMLIGTPPQKLQILVDTGSSNFAVAGTPHSYIDT-----YFDTERSSTYRSKGFDV 145
Qу
            79 YYGDIGIGTPPQCFTVVFDTGSSNLWVPSIHCKILDIACWVHHKYNSDKSSTYVKNGTSF 138
Db
        146 TVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIAT---IFESENFFLPGI----KWNGIL 197
Qу
            139 DIHYGSGSLSGYLSQDTVSVPCKSDQSKARGIKVEKQIF-GEATKQPGIVFVAAKFDGIL 197
Db
        198 GLAYATLAKPSSSLETFFDSLVTQANI-PNVFSMQMCGAGLPVAGSGTNGGSLVLGGIEP 256
Qy
            198 GMGYPHIS--VNNVLPVFDNLMQQKLVDKNIFSFY----LNRDPEGQPGGELMLGGTDS 250
Db
        257 SLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVF 316
Qy
             | |:: | : : |:|: : :||:| : | | |:
                                               :|||:||:||
        251 KYYHGELSYLNVTRKAYWQVHMDQLEVGNE-LTL-CK--GGCEAIVDTGTSLLVGPVEEV 306
Db
        317 DAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITIL 376
Qу
                                                  : |
             ::1: 11
        307 KELOKAIGAVPLI------QGEYMIPCEKVSSL 333
Db
        377 PQLYIQPMMGAGLNYEC----YRFGIS-----PSTNALVIGATVMEG 414
Qy
            334 PTVYLK--LG-GKNYELHPDKYILKVSQGGKTICLSGFMGMDIPPPSGPLWILGDVFIGS 390
Db
        415 FYVIFDRAQKRVGFA 429
Qy
            : | : | | | | | | | |
        391 YYTVFDRDNNRVGFA 405
Db
```

Search completed: March 4, 2004, 15:40:59
Job time: 29.1043 secs

# GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: March 4, 2004, 15:30:05; Search time 28.1043 Seconds

(without alignments)

1772.942 Million cell updates/sec

Title: US-09-668-314C-2

Perfect score: 2687

Sequence: 1 MGALARALLLPLLAQWLLRA.....RPRDPEVVNDESSLVRHRWK 518

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 283366 seqs, 96191526 residues

Total number of hits satisfying chosen parameters: 283366

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database: PIR 78:\*

1: pir1:\*

2: pir2:\*

3: pir3:\*

4: pir4:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

		8				
Result		Query				
No.	Score	Match	Length	DB	ID	Description
1	1178.5	43.9	501	2	A59090	aspartic proteinas
2	367.5	13.7	383	2	JC7573	pepsinogen C - Afr
3	363.5	13.5	377	1	PEMQCJ	gastricsin (EC 3.4
4	355.5	13.2	384	2	A39314	gastricsin (EC 3.4
5	355	13.2	389	2	JE0371	pepsin C (EC 3.4.2
6	353	13.1	388	2	A29937	gastricsin (EC 3.4
7	351.5	13.1	388	2	JC7246	pepsinogen C - com
8	324.5	12.1	394	2	B43356	gastricsin (EC 3.4
9	320	11.9	385	2	JC7575	pepsinogen A - bul
10	320	11.9	402	1	REMSK	renin (EC 3.4.23.1
11	313.5	11.7	509	2	S66516	oryzasin (EC 3.4.2
12	313	11.6	392	1	A24608	gastricsin (EC 3.4
13	310	11.5	383	2	A41443	pepsin (EC 3.4.23.

14	308.5	11.5	412	1	KHHUD	cathepsin D (EC 3.
15	306.5	11.4	410	1	KHMSD	cathepsin D (EC 3.
16	305.5	11.4	401	1	REMSS	renin (EC 3.4.23.1
17	305	11.4	384	2	JC7574	pepsinogen A - Afr
18	305	11.4	407	1	KHRTD	cathepsin D (EC 3.
19	302	11.2	405	2	A25379	saccharopepsin (EC
20	301.5	11.2	398	2	S66465	cathepsin E (EC 3.
21	300.5	11.2	387	2	C38302	pepsin (EC 3.4.23.
22	299	11.1	398	2	I51185	cathepsin D (EC 3.
23	298.5	11.1	387	2	D38302	pepsin (EC 3.4.23.
24	298.5	11.1	400	2	I47099	renin (EC 3.4.23.1
25	297	11.1	388	1	PEHU	pepsin A (EC 3.4.2
26	296	11.0	388	2	A30142	pepsin A (EC 3.4.2
27	296	11.0	388	2	B30142	pepsin A (EC 3.4.2
28	294.5	11.0	388	1	S19684	pepsin A (EC 3.4.2
29	292	10.9	506	2	T07915	probable aspartic
30	291	10.8	388	1	S19682	pepsin A (EC 3.4.2
31	291	10.8	402	1	RERTK	renin (EC 3.4.23.1
32	291	10.8	406	1	REHUK	renin (EC 3.4.23.1
33	290.5	10.8	396	2	S36865	cathepsin E (EC 3.
34	289	10.8	387	2	E38302	pepsin (EC 3.4.23.
35	288	10.7	387	2	B38302	pepsin (EC 3.4.23.
36	288	10.7	388	1	PEMQAJ	pepsin A (EC 3.4.2
37	287.5	10.7	632	2	T45858	hypothetical prote
38	287	10.7	391	2	A43356	cathepsin E (EC 3.
39	287	10.7	396	2	A34401	cathepsin E (EC 3.
40	286.5	10.7	334	2	JC4870	pepsin A (EC 3.4.2
41	286	10.6	382	1	PECH	pepsin A (EC 3.4.2
42	286	10.6	388	1	PEMQAR	pepsin A (EC 3.4.2
43	285.5	10.6	387	2	JC7245	pepsinogen A - com
44	285	10.6	396	2	T47207	aspartic proteinas
45	284.5	10.6	386	1	PEPG	pepsin A (EC 3.4.2

#### ALIGNMENTS

```
RESULT 1
A59090
aspartic proteinase (EC 3.4.23.-) BACE precursor - human
N;Alternate names: beta-secretase; beta-site APP cleaving enzyme
C;Species: Homo sapiens (man)
C;Date: 29-Oct-1999 #sequence_revision 29-Oct-1999 #text_change 11-May-2000
C;Accession: A59090
R;Vassar, R.; Bennett, B.D.; Babu-Khan, S.; Kahn, S.; Mendiaz, E.A.; Denis, P.;
Teplow, D.B.; Ross, S.; Amarante, P.; Loeloff, R.; Luo, Y.; Fisher, S.; Fuller,
J.; Edenson, S.; Lile, J.; Jarosinski, M.A.; Biere, A.L.; Curran, E.; Burgess,
T.; Louis, J.C.; Collins, F.; Treanor, J.; Rogers, G.; Citron, M.
Science 286, 735-741, 1999
```

A; Title: beta-Secretase cleavage of Alzheimer's amyloid precursor protein by the transmembrane aspartic protease BACE.

A; Reference number: A59090; MUID: 20002972; PMID: 10531052

A; Note: submitted to GenBank, September 1999

A; Accession: A59090

A; Status: not compared with conceptual translation

A; Molecule type: mRNA A; Residues: 1-501 < VAS>

```
A;Cross-references: GB:AF190725; NID:q6118538; PIDN:AAF04142.1; PID:q6118539
C; Genetics:
A; Gene: BACE
C; Superfamily: beta-secretase
C; Keywords: Alzheimer's disease; aspartic proteinase; brain; glycoprotein;
hydrolase; protein digestion; transmembrane protein; zymogen
F;1-21/Domain: signal sequence #status predicted <SIG>
F;22-45/Domain: propeptide #status predicted <PRO>
F;46-501/Product: acid proteinase BACE #status predicted <MAT>
F;461-477/Domain: transmembrane #status predicted <TRN>
F;93,289/Active site: Asp #status predicted
F;153,172,223,354/Binding site: carbohydrate (Asn) (covalent) #status predicted
F:330-380/Disulfide bonds: #status predicted
                       43.9%; Score 1178.5; DB 2; Length 501; 46.2%; Pred. No. 3.8e-80;
 Query Match
 Best Local Similarity
 Matches 240; Conservative 82; Mismatches 164; Indels
                                                          33; Gaps
                                                                      9;
           7 ALLLPLLAOWLLRAAPELAPAPFT----LPLRVAAATNRVVAPTPGPGTPAERHADGLA 61
Qу
                           : || |
                                      1111
                                                        1 11 1 1:
           2 AQALPWLLLWM---GAGVLPAHGTQHGIRLPLRSGLG------GAPL----GLR 42
Db
          62 LALE--PALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVA 119
Qу
            | 1
                      - 1
                           :|: |||||:| ||:||:|| :|:||| | | ||:|||
          43 LPRETDEEPEEPGRRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFAVG 102
Db
         120 GTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIAT 179
Qy
              11:: 1: : []]
                                   103 AAPHPFLHRYYQRQLSSTYRDLRKGVYVPYTQGKWEGELGTDLVSIPHGPNVTVRANIAA 162
Db
         180 IFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPV 239
Qу
             163 ITESDKFFINGSNWEGILGLAYAEIARPDDSLEPFFDSLVKQTHVPNLFSLQLCGAGFPL 222
Db
         240 AGS---GTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYN 296
Qy
                  223 NOSEVLASVGGSMIIGGIDHSLYTGSLWYTPIRREWYYEVIIVRVEINGQDLKMDCKEYN 282
Db
         297 ADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFP 356
QУ
              111: 11
         283 YDKSIVDSGTTNLRLPKKVFEAAVKSIKAASSTEKFPDGFWLGEQLVCWQAGTTPWNIFP 342
Db
         357 KISIYLRDENSSRSFRITILPQLYIQPMMGAGLNY-ECYRFGISPSTNALVIGATVMEGF 415
Qy
             : :||:| || |:
         343 VISLYLMGEVTNOSFRITILPQQYLRPVEDVATSQDDCYKFAISQSSTGTVMGAVIMEGF 402
Db
         416 YVIFDRAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYAL 475
Qγ
                                   :: ||| | | |:
                                                              ::|:
             ||:||:||:||:||
                                                           403 YVVFDRARKRIGFAVSACHVHDEFRTAAVEGPFVTLDMEDCGYNIPOTDESTLMTIAYVM 462
Db
         476 MSVCGAILLVLIVLLLLPFRCQR--RPRDPEVVNDESSL 512
Qy
              ::| |: :: : |:: :|| | | | | | | |
         463 AAIC-ALFMLPLCLMVCQWRCLRCLRQQHDDFADDISLL 500
Db
```

```
pepsinogen C - African clawed frog
N; Alternate names: progastricsin
C; Species: Xenopus laevis (African clawed frog)
C; Date: 30-Jun-2001 #sequence revision 30-Jun-2001 #text change 03-Aug-2001
C; Accession: JC7573; PC7118
R; Ikuzawa, M.; Inokuchi, T.; Kobayashi, K.; Yasumasu, S.
J. Biochem. 129, 147-153, 2001
A; Title: Amphibian pepsinogens: Purification and characterization of Xenopus
pepsinogens, and molecular cloning of Xenopus and bullfrog pepsinogens.
A; Reference number: JC7573; MUID:21064922; PMID:11134969
A; Contents: Stomach
A; Accession: JC7573
A; Molecule type: mRNA
A; Residues: 1-383 <IKU>
A; Cross-references: DDBJ:AB045379
A; Accession: PC7118
A; Molecule type: protein
A; Residues: 17-68 <IK2>
C; Comment: This protein is a zymogen for gastric aspartic proteinase, with
pepsin-like activity.
C; Genetics:
A; Gene: PgC
C; Superfamily: pepsin
C; Keywords: stomach; zymogen
                       13.7%; Score 367.5; DB 2; Length 383;
 Query Match
 Best Local Similarity 28.9%; Pred. No. 9.1e-20;
                                                                   25;
 Matches 132; Conservative 70; Mismatches 154; Indels 101; Gaps
          1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qу
            1 | | | : :: :: | : : : : |
                                           1: || 1 |
          1 MKFLILALVCLQLSEGIIR-----VPLKKFKSMREVMRENGIKAPLVDPAT----KYYNQY 52
Db
         61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qy
                                       :|::|
         53 ATAYEP----LSNYMDM-----SYYGEISIGTPPQNFLVLFDTGSSNLWVAS 95
Db
         121 TPHSYIDT-----YFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSF 173
Qу
            1 :
                         96 T---YCQSQACTNHPLFNPSQSSTYSSNQQQFSLQYGTGSLTGILGYDTVTIQ----- 145
Db
         174 LVNIATIFESENFFL---PG----IKWNGILGLAYATLAKPSSSLETFFDSLVTQANI 224
Qy
                    : | | | |
                                   :::||||||| ::| :: |
         146 --NVA--ISQQEFGLSETEPGTNFVYAQFDGILGLAYPSIAVGGAT--TVMQGMM-QQNL 198
Db
         225 PN--VFSMOMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLE 282
Qу
             199 LNQPIFGFYLSGQ-----SSQNGGEVAFGGVDQNYYTGQIYWTPVTSETYWQIGIQGFS 252
Db
         283 IGGOSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQL 342
Qу
            253 INGQATGW-CSQ--GCQAIVDTGTSLLTAPQSVFSSLIQSIG-----AQQDQNGQYV 301
Db
         343 ACWTNSETPWSYFPKISIYLRDENSSRSFRITILPQLYI-QPMMGAGLNYECYRFGIS-- 399
Qу
               302 VSCSNIQN----LPTISFTI----SGVSFPLP--PSAYVLQQSSG-----YC-TIGIMPT 345
Db
```

```
Qу
          400 -- PSTNA---LVIGATVMEGFYVIFDRAQKRVGFAAS 431
                         ::1
                              : :| ::|
                                             : | | | | :
                \mathbf{I} \mathbf{I} \mathbf{I}
          346 YLPSQNGQPLWILGDVFLREYYSVYDLGNNQVGFATA 382
Db
RESULT 3
PEMOCJ
qastricsin (EC 3.4.23.3) precursor - Japanese macaque (fragment)
N; Alternate names: pepsin C
C; Species: Macaca fuscata (Japanese macaque)
C;Date: 13-Aug-1986 #sequence revision 19-Oct-1995 #text change 18-Jun-1999
C; Accession: S19683; A00986; A22402; S16066
R; Kageyama, T.; Tanabe, K.; Koiwai, O.
Eur. J. Biochem. 202, 205-215, 1991
A; Title: Development-dependent expression of isozymogens of monkey pepsinogens
and structural differences between them.
A: Reference number: S19681; MUID: 92037645; PMID: 1935977
A; Accession: S19683
A; Molecule type: mRNA
A; Residues: 1-377 <KAG>
A; Cross-references: EMBL: X59754; NID: q38072; PIDN: CAA42426.1; PID: q38073
R; Kageyama, T.; Takahashi, K.
J. Biol. Chem. 261, 4406-4419, 1986
A; Title: The complete amino acid sequence of monkey progastricsin.
A; Reference number: A00986; MUID: 86168133; PMID: 3514597
A; Accession: A00986
A; Molecule type: protein
A; Residues: 6-330, 'V', 332-349, 'VY', 350-377 <KA2>
R; Kageyama, T.; Takahashi, K.
J. Biochem. 97, 1235-1246, 1985
A; Title: Monkey pepsinogens and pepsins. VII. Analysis of the activation process
and determination of the NH2-terminal 60-residue sequence of Japanese monkey
progastricsin, and molecular evolution of pepsinogens.
A; Reference number: A22402; MUID: 85289106; PMID: 3928607
A; Accession: A22402
A; Molecule type: protein
A; Residues: 6-65 < KA3>
C; Comment: This enzyme has more restricted specificity than pepsin A.
C; Comment: The enzyme is activated in a two-step process that gives rise to two
end products. The shorter, Ser-gastricsin, is the major product.
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; gastric juice; hydrolase; protein digestion;
stomach
F;1-5/Domain: signal sequence (fragment) #status predicted <SIG>
F;6-377/Product: progastricsin #status experimental <ZYM>
F;6-45/Domain: activation peptide #status experimental <APT>
F;46-377/Product: Gly-gastricsin #status experimental <MIN>
F;49-377/Product: Ser-gastricsin #status experimental <MAT>
F;31-32/Cleavage site: Phe-Leu (pepsin) #status experimental
F;45-46/Cleavage site: Phe-Gly (pepsin) #status experimental
F;48-49/Cleavage site: Leu-Ser (pepsin) #status experimental
F;80,265/Active site: Asp #status predicted
F;93-98,256-260,299-332/Disulfide bonds: #status experimental
                           13.5%; Score 363.5; DB 1; Length 377;
  Query Match
  Best Local Similarity 28.9%; Pred. No. 1.8e-19;
                               65; Mismatches 118; Indels 107; Gaps
                                                                              19;
  Matches 118; Conservative
```

```
56 HADGLALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSN 115
Qу
           44 HFGDLSVSYEP-----MAYMD-----AAYFGEISIGTPPONFLVLFDTGSSN 85
Db
        116 FAV-----AGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPK 167
Qу
             86 LWVPSVYCQSQACTSHS----RFNPSESSTYSTNGQTFSLQYGSGSLTGFFGYDTLTV-- 139
Db
        168 GFNTSFLVNIATIFESENFFLPG----IKWNGILGLAYATLAKPSSSLETFFDSLVTQA 222
Qу
                        140 ---QSIQVPNQEFGLSEN--EPGTNFVYAQFDGIMGLAYPTLSVDGAT--TAMQGMVQEG 192
Db
        223 NIPN-VFSMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKL 281
Qy
                             |::||::| ||:: ||| | |:: |: || |:|| |:
             : : :||: :
        193 ALTSPIFSVYLSDQ-----QGSSGGAVVFGGVDSSLYTGQIYWAPVTQELYWQIGIEEF 246
Db
        282 EIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQ 341
Qγ
                          :||||:||:||:||
                   1 1
        247 LIGGQASGW-CSE--GCQAIVDTGTSLLTVPQQYMSALLQA-----TGAQ 288
Db
        342 LACWTNSETPWSYF-----PKISIYLRDENSSRSFRITILPQLYIQPMMGAGLNY 391
Qу
                 289 -----EDEYGQFLVNCNSIQNLPTLTFII-----NGVEFPLPPSSYI-----LNN 328
Db
        392 ECY-RFGISP-----STNALVIGATVMEGFYVIFDRAOKRVGFAAS 431
Qу
             | |:| | ::| ::|::|:
        329 NGYCTVGVEPTYLSAQNSQPLWILGDVFLRSYYSVYDLSNNRVGFATA 376
Db
RESULT 4
A39314
gastricsin (EC 3.4.23.3) precursor - bullfrog
C; Species: Rana catesbeiana (bullfrog)
C;Date: 19-Jun-1992 #sequence revision 19-Jun-1992 #text change 22-Jun-1999
C; Accession: A39314
R; Yakabe, E.; Tanji, M.; Ichinose, M.; Goto, S.; Miki, K.; Kurokawa, K.; Ito,
H.; Kageyama, T.; Takahashi, K.
J. Biol. Chem. 266, 22436-22443, 1991
A; Title: Purification, characterization, and amino acid sequences of pepsinogens
and pepsins from the esophageal mucosa of bullfrog (Rana catesbeiana).
A; Reference number: A39314; MUID: 92042186; PMID: 1939266
A; Accession: A39314
A; Status: preliminary
A; Molecule type: mRNA
A; Residues: 1-384 <YAK>
A;Cross-references: GB:M73750; NID:g213687; PIDN:AAA49530.1; PID:g213688
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; hydrolase; protein digestion
                      13.2%; Score 355.5; DB 2; Length 384;
 Query Match
 Best Local Similarity 26.5%; Pred. No. 7.2e-19;
 Matches 120; Conservative 73; Mismatches 136; Indels 123; Gaps
                                                                  21;
         23 ELAPAPFTLPLRVAAATNRVV-----APTPGPGTPAERHADGLALALEPALASPAGAAN 76
Qy
            12 QLSEGIIKVPLKKFKSMREVMRDHGIKAPVVDPAT---KYYNNFATAFEP-----LAN 61
Db
```

```
77 FLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGTPHSYIDT-----Y 129
Qу
           Db
         62 YMDM-----SYYGEISIGTPPQNFLVLFDTGSSNLWV---PSTYCQSQACTNHPQ 108
        130 FDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFESENFFL- 188
QУ
           109 FNPSQSSSYSSNQQQFSLQYGTGSLTGILGYDTVQIQ-----NIA--ISQQEFGLS 157
Db
        189 ---PG----IKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPN--VFSMQMCGAGLP 238
QУ
                   :::|||||||||:|: :: | :: | |: | :|: : |
Db
        158 VTEPGTNFVYAQFDGILGLAYPSIAEGGAT--TVMQGMI-QQNLINQPLFAFYLSG---- 210
        239 VAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNAD 298
Qν
              Db
        211 -QQNSQNGGEVAFGGVDQNYYSGQIYWTPVTSETYWQIGIQGFSVNGQATGW-CSQ--GC 266
        299 KAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQ-----LACWTNSET 350
Qy
           267 OGIVDTGTSLLTAPQSVFSSLMQSI------GAQQDQNGQYAVSCSNIQSL 311
Db
        351 PWSYFP----KISIYLRDENS---SRSFRITILPQLYIQPMMGAGLNYECYRFGIS 399
Qy
           312 PTISFTISGVSFPLPPSAYVLQQNSGYCTIGIMPTYLPSQNGQPLW----- 357
Db
Qу
        400 PSTNALVIGATVMEGFYVIFDRAQKRVGFAAS 431
            ::| : :| ::| :|||::
Dh
        358 -----ILGDVFLRQYYSVYDLGNNQVGFAAA 383
RESULT 5
JE0371
pepsin C (EC 3.4.23.-) precursor - chicken
N; Alternate names: pepsinogen C
C; Species: Gallus gallus (chicken)
C;Date: 23-Jul-1999 #sequence revision 23-Jul-1999 #text change 11-May-2000
C; Accession: JE0371
R; Sakamoto, N.; Saiga, H.; Yasugi, S.
Biochem. Biophys. Res. Commun. 250, 420-424, 1998
A; Title: Analysis of temporal expression pattern and cis-regulatory sequences of
chicken pepsinogen A and C.
A; Reference number: JE0370; MUID: 98440813; PMID: 9753645
A; Accession: JE0371
A; Status: preliminary
A; Molecule type: mRNA
A; Residues: 1-389 <SAK>
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; hydrolase
                     13.2%; Score 355; DB 2; Length 389;
 Query Match
 Best Local Similarity 28.7%; Pred. No. 7.9e-19;
 Matches 114; Conservative 58; Mismatches 121; Indels 104; Gaps
         75 ANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGT-----PHSYI 126
Qу
           56 SNFATAYEPLANNMDMSYYGEISIGTPPQNFLVLFDTGSSNLWVPSTLCQSQACANHN-- 113
Db
```

```
127 DTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFN-----TS 172
Qу
                                :::| || || || || || || :| :
                     111: ::
         114 --EFDPNESSTFSTQDEFFSLQYGSGSLTGIFGFDTVTI-QGISITNQEFGLSETEPGTS 170
Db
         173 FLVNIATIFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPN--VFSM 230
Qу
                                 ::||||||: :: |: |
                                                          : | |: : |||
              ||:
Db
         171 FLYS-----PFDGILGLAFPSI---SAGGATTVMQKMLQENLLDFPVFSF 212
         231 QMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNL 290
Qу
                         |: || || || ||::|:|| || ||:||: ||:|| ||
         213 YLSGQ-----EGSQGGELVFGGVDPNLYTGQITWTPVTQTTYWQIGIEDFAVGGQSSGW 266
Db
         291 DCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSET 350
Qу
                   : |||:||:|| :| :| :: : : : |
                                                           :| :|
         267 -CSQ--GCQGIVDTGTSLLTVPNQVFTELMQYIG----AQADD---SGQYVASCSNIE- 314
Db
         351 PWSYFPKI-----SIYLRDENS---SRSFRITILPQLYIQPMMGAGLNYECY 394
Qy
                                  ||:
         315 ---YMPTITFVISGTSFPLPPSAYMLQSNSDYCTVGIESTYLPSQTGQPLW----- 362
Db
         395 RFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qу
                        ::| : :| |:|
                                          : | | | | :
         363 -----ILGDVFLRVYYSIYDMGNNQVGFATA 388
Db
RESULT 6
A29937
qastricsin (EC 3.4.23.3) precursor - human
N; Alternate names: pepsin C; pepsinogen C
C; Species: Homo sapiens (man)
C;Date: 17-Oct-1988 #sequence revision 17-Oct-1988 #text change 31-Mar-2000
C; Accession: A29937; A31811; PX0028; I54213; A91125; A23458
R; Hayano, T.; Sogawa, K.; Ichihara, Y.; Fujii-Kuriyama, Y.; Takahashi, K.
J. Biol. Chem. 263, 1382-1385, 1988
A; Title: Primary structure of human pepsinogen C gene.
A; Reference number: A29937; MUID: 88087276; PMID: 3335549
A; Accession: A29937
A; Molecule type: DNA
A; Residues: 1-388 <HAY>
R; Taggart, R.T.; Cass, L.G.; Mohandas, T.K.; Derby, P.; Barr, P.J.; Pals, G.;
Bell, G.I.
J. Biol. Chem. 264, 375-379, 1989
A; Title: Human pepsinogen C (progastricsin). Isolation of cDNA clones,
localization to chromosome 6, and sequence homology with pepsinogen A.
A; Reference number: A31811; MUID: 89079679; PMID: 2909526
A; Accession: A31811
A; Molecule type: mRNA
A; Residues: 1-388 <TAG>
A;Cross-references: GB:J04443; NID:q551175; PIDN:AAA60074.1; PID:q551176
R; Athauda, S.B.P.; Tanji, M.; Kageyama, T.; Takahashi, K.
J. Biochem. 106, 920-927, 1989
A; Title: A comparative study on the NH2-terminal amino acid sequences and some
other properties of six isozymic forms of human pepsinogens and pepsins.
A; Reference number: PX0023; MUID: 90130402; PMID: 2515193
A; Accession: PX0028
A; Molecule type: protein
A; Residues: 17-101 <ATH>
```

```
R; Pals, G.; Azuma, T.; Mohandas, T.K.; Bell, G.I.; Bacon, J.; Samloff, I.M.;
Walz, D.A.; Barr, P.J.; Taggart, R.T.
Genomics 4, 137-148, 1989
A; Title: Human pepsinogen C (progastricsin) polymorphism: evidence for a single
locus located at 6p21.1-pter.
A; Reference number: I54213; MUID:89290840; PMID:2567697
A; Accession: I54213
A; Status: translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-388 < RES>
A;Cross-references: GB:M23077; NID:g189830; PIDN:AAA60063.1; PID:g387015;
GB: J03063
A; Note: parts of this sequence, including the amino end and carboxyl ends of the
mature protein, were determined by protein sequencing
R; Foltmann, B.; Jensen, A.L.
Eur. J. Biochem. 128, 63-70, 1982
A; Title: Human progastricsin. Analysis of intermediates during activation into
gastricsin and determination of the amino acid sequence of the propart.
A; Reference number: A91125; MUID: 83079318; PMID: 6816595
A; Accession: A91125
A; Molecule type: protein
A; Residues: 17-39, 'ED', 42-51, 'S', 53-64 <FOL>
A; Note: pro-form; 29-Leu was also found
A; Note: activation at pH 2 is proposed to involve conformation change, cleavage
after Phe-42, and cleavage after Leu-59
C; Genetics:
A; Gene: GDB: PGC
A; Cross-references: GDB:119485; OMIM:169740
A; Map position: 6p21.3-6p21.1
A; Introns: 20/2; 70/3; 110/1; 149/3; 216/2; 256/2; 305/3; 338/3
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; hydrolase; protein digestion; stomach; zymogen
F;1-16/Domain: signal sequence #status predicted <SIG>
F;17-59/Domain: propeptide #status experimental <PRO>
F;60-388/Product: gastricsin #status experimental <MAT>
                         13.1%; Score 353; DB 2; Length 388;
  Query Match
  Best Local Similarity 29.1%; Pred. No. 1.1e-18;
 Matches 120; Conservative 65; Mismatches 120; Indels 108; Gaps
                                                                           21;
           52 PAERHADG-LALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVD 110
Qy
              || :: | |:: ||
                                        :|:|
                                                      1: 1: |||||
           50 PAWKYRFGDLSVTYEP-----MAYMD-----AAYFGEISIGTPPQNFLVLFD 91
Db
          111 TGSSNFAV-----AGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDL 162
Qу
                                             \Box\Box\Box
              11111 1
                             1 1 11
                                        |:
                                                        :::| || || || |
           92 TGSSNLWVPSVYCQSQACTSHS----RFNPSESSTYSTNGQTFSLQYGSGSLTGFFGYDT 147
Db
          163 VTIPKGFNTSFLVNIATIFESENFFLPG----IKWNGILGLAYATLAKPSSSLETFFDS 217
Qу
                                               :::||:||||
                                 111
                                      - 11
                                                                ::
                      1 1
          148 LTV----QSIQVPNQEFGLSEN--EPGTNFVYAQFDGIMGLAYPALSVDEAT--TAMQG 198
Db
         218 LVTQANIPN-VFSMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQI 276
Qу
                                       :|::::|||:::
          199 MVQEGALTSPVFSVYLSNQ-----QGSSGGAVVFGGVDSSLYTGQIYWAPVTQELYWQI 252
Db
          277 EILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGF 336
Qу
```

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| : ||||: | | :|||:||:||:||:||
Db
         253 GIEEFLIGGQASGW-CSE--GCQAIVDTGTSLLTVPQQYMSALLQA----- 295
         337 WTGSOLACWTNSETPWSYF-----PKISIYLRDENSSRSFRITILPQLYIQPMMG 386
Qу
                                 | :: :
                      | : |
                                                         : | ||
         296 -TGAQ-----EDEYGQFLVNCNSIQNLPSLTFII-----NGVEFPLPPSSYI---- 336
Db
         387 AGLNYECY-RFGISP----STNA---LVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qу
              1: 1: 1: 1
                                    ::| : :| ::|
         337 -- LSNNGYCTVGVEPTYLSSQNGQPLWILGDVFLRSYYSVYDLGNNRVGFATA 387
Db
RESULT 7
JC7246
pepsinogen C - common marmoset
C; Species: Callithrix jacchus (common marmoset)
C;Date: 09-Jun-2000 #sequence revision 09-Jun-2000 #text change 21-Jul-2000
C:Accession: JC7246
R; Kaqeyama, T.
J. Biochem. 127, 761-770, 2000
A; Title: New world monkey pepsinogens A and C, and prochymosins. Purification,
characterization of enzymatic properties, cDNA cloning, and molecular evolution.
A; Reference number: JC7245
A; Accession: JC7246
A; Molecule type: mRNA
A; Residues: 1-388 < KAG>
A; Cross-references: DDBJ:AB038385
A; Experimental source: strain NW791
C; Comment: This protein, a zymogen of pepsins, is the major proteolytic enzyme
in vertebrate gastric juices. It plays roles in gastric digestion, and is a
useful molecular marker for clarifying the evolution of mammalian orders and
families.
C; Superfamily: pepsin
C; Keywords: gastric juice; zymogen
                        13.1%; Score 351.5; DB 2; Length 388;
 Query Match
 Best Local Similarity 30.1%; Pred. No. 1.4e-18;
 Matches 112; Conservative 56; Mismatches 115; Indels
                                                           89;
                                                                Gaps
                                                                      17;
          92 YYLEMLIGTPPQKLQILVDTGSSNFAV-----AGTPHSYIDTYFDTERSSTYRSKGF 143
Qу
             1: 1: 111111 :1 111111 1
                                             1 1 11
                                                      |: |||| | |
          73 YFGEISIGTPPQNFLVLFDTGSSNLWVPSVYCQSQACTSHS----RFNPSASSTYSSNGQ 128
Db
         144 DVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFESENFFLPG----IKWNGILG 198
Qy
               :::| || || || || || ||
                                               1 1
         129 TFSLOYGSGSLTGFFGYDTLTV----QSIQVPNQEFGLSEN--EPGTNFVYAQFDGIMG 181
Db
         199 LAYATLAKPSSSLETFFDSLVTQANIPN-VFSMQMCGAGLPVAGSGTNGGSLVLGGIEPS 257
Qy
                    :: | :: : : | | | :
                                                     1::||::: ||:: |
         182 LAYPALSMGGAT--TAMQGMLQEGALTSPVFSFYLSNQ-----QGSSGGAVIFGGVDSS 233
Db
         258 LYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFD 317
Οv
             234 LYTGOIYWAPVTOELYWQIGIEEFLIGGQASGW-CSE--GCQAIVDTGTSLLTVPQQYMS 290
Db
         318 AVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYF----PKISIYLRDENS 367
Qу
                               11:1 : 1
             1:11
```

```
291 AFLEA-----TGAQ-----EDEYGOFLVNCDSIONLPTLTFII---- 323
Db
          368 SRSFRITILPOLYIOPMMGAGLNYECY-RFGISP----STNALVIGATVMEGFYVIF 419
Qv
                    : 1 11
                                1: | 1: |
                                                  i
                                                      ::1
                                                           : :1:1
Db
          324 -NGVEFPLPPSSYI-----LSNNGYCTVGVEPTYLSSQNSQPLWILGDVFLRSYYSVF 375
          420 DRAQKRVGFAAS 431
Qу
                 11111:
             1
          376 DLGNNRVGFATA 387
Db
RESULT 8
B43356
gastricsin (EC 3.4.23.3) precursor - guinea pig
. N; Alternate names: pepsin C
C; Species: Cavia porcellus (guinea pig)
C;Date: 03-Feb-1994 #sequence revision 03-Feb-1994 #text change 22-Jun-1999
C:Accession: B43356
R; Kageyama, T.; Ichinose, M.; Tsukada, S.; Miki, K.; Kurokawa, K.; Koiwai, O.;
Tanji, M.; Yakabe, E.; Athauda, S.B.; Takahashi, K.
J. Biol. Chem. 267, 16450-16459, 1992
A; Title: Gastric procathepsin E and progastricsin from guinea pig. Purification,
molecular cloning of cDNAs, and characterization of enzymatic properties, with
special reference to procathepsin E.
A; Reference number: A43356; MUID: 92355614; PMID: 1644829
A; Accession: B43356
A; Status: preliminary
A; Molecule type: mRNA
A; Residues: 1-394 <KAG>
A;Cross-references: GB:M88652; NID:q191296; PIDN:AAA37053.1; PID:q191297
A; Note: sequence extracted from NCBI backbone (NCBIN:110805, NCBIP:110806)
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; gastric juice; hydrolase; protein digestion;
stomach
  Query Match
                        12.1%; Score 324.5; DB 2; Length 394;
  Best Local Similarity 29.0%; Pred. No. 1.5e-16;
  Matches 107; Conservative 63; Mismatches 116; Indels
                                                           83; Gaps
                                                                      18;
          92 YYLEMLIGTPPQKLQILVDTGSSNF-----AVAGTPHSYIDTYFDTERSSTYRSKGF 143
Qу
             1: :: :||||| |:| |||||
                                           ::| | |
                                                     1 1:
          79 YFGQISLGTPPQSFQVLFDTGSSNLWVPSVYCSSLACTTH----TRFNPRDSSTYVATDQ 134
Db
          144 DVTVKYTQGSWTGFVGEDLVTI----PK-GFNTSFLVNIATIFESENFFLPG----IK 192
Qy
               1:1 11
          135 SFSLEYGTGSLTGVFGYDTMTIQDIQVPKQEFGLS-----ETE----PGSDFVYAE 181
Db
          193 WNGILGLAYATLAKPSSSLETFFDSLVTQANI-PNVFSMQMCGAGLPVAGS--GTNGGSL 249
Qу
             182 FDGILGLGYPGLSEGGAT--TAMQGLLREGALSQSLFSVYL-----GSQQGSDEGQL 231
Db
          250 VLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLL 309
Οv
             232 ILGGVDESLYTGDIYWTPVTQELYWQIGIEGFLIDGSASGWCSR---GCQGIVDTGTSLL 288
 Db
          310 RLPOKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSR 369
QУ
                     :|:|: |::| ::| ::| ::
              : |
```

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Db
         289 TVPSDYLSTLVOAIGAEE--NEYGEYF-----VSCSSIODLPTLTFVISGV------ 332
         370 SFRITILPOLYIOP----MMGAGLNYECYRFGISPSTN--ALVIGATVMEGFYVIFDRA 422
Qу
                               1:1
                                     - 1
                                            :11
                                                      ::1
                                                          : :|::||
Db
         333 --EFPLSPSAYILSGENYCMVGLESTY-----VSPGGGEPVWILGDVFLRSYYSVYDLA 384
Qу
         423 QKRVGFAAS 431
               11111:
         385 NNRVGFATA 393
Db
RESULT 9
JC7575
pepsinogen A - bullfrog
C; Species: Rana catesbeiana (bullfrog)
C;Date: 30-Jun-2001 #sequence revision 30-Jun-2001 #text change 03-Aug-2001
C; Accession: JC7575
R; Ikuzawa, M.; Inokuchi, T.; Kobayashi, K.; Yasumasu, S.
J. Biochem. 129, 147-153, 2001
A; Title: Amphibian pepsinogens: Purification and characterization of Xenopus
pepsinogens, and molecular cloning of Xenopus and bullfrog pepsinogens.
A; Reference number: JC7573; MUID:21064922; PMID:11134969
A; Contents: Stomach
A: Accession: JC7575
A; Molecule type: mRNA
A; Residues: 1-385 <IKU>
A; Cross-references: DDBJ:AB045376
C; Comment: This protein is a zymogen for gastric aspartic proteinase, with
pepsin-like activity.
C; Genetics:
A; Gene: PqA
C; Superfamily: pepsin
C; Keywords: stomach; zymogen
                        11.9%; Score 320; DB 2; Length 385;
 Query Match
 Best Local Similarity 27.8%; Pred. No. 3.2e-16;
 Matches 111; Conservative 67; Mismatches 147; Indels
                                                                         15;
          50 GTPAERHADGLALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILV 109
Qy
             | ::| | |:|| :| :||
                                                     1: : |||||
          39 GDYLKKHHYNPATKYFPSLAQASG-----EPLQNYMDIEYFGTISIGTPPQSFTVIF 90
Db
         110 DTGSSNFAVAGTPHSYIDT-----YFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDL 162
Qу
                                      |: ::|||:::
             1:::1 11 :11:1 1
Db
          91 DTGSSNLWV---PSVYCSSPACTNHHMFNPQQSSTFQATNTPVSIQYGTGSMSGFLGYDT 147
         163 VTIPKGFNTSFLVNIATIFESE-NFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQ 221
Qу
                         - 1
                                                                ||:: |
Db
         148 VOVG---NIOITNOIFGLSOSEPGSFLYYSPFDGILGLAFPSLA--SSOATPVFDNMWNO 202
         222 ANIP-NVFSMOMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYOIEILK 280
Qy
                              -: |:| :: ||:: | | |:: : |: | |:|| :
         203 GLIPODLFSVYL-----SSQGQSGSFVLFGGVDTSYYTGNLNWVPLTAETYWQITVDS 255
Db
         281 LEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGS 340
Qу
                           : ||||:||:||
                                                | ::
                                                       1:
                                                            : |:| :
Db
         256 ISIGGOVIACS----GSCSAIVDTGTSLLAGPSTPI-ANIOYYIGAN---ODSNGOYV-- 305
```

```
341 QLACWTNSETPWSYFP-----KISIYLRDENSS-RSFRITILPQLYIQPMMGAGLN 390
Qу
                   1 1 1
                                        | |:|
                                                - 1
                                                      1:
          306 -INCNNISNMPTVVFTINGVQYPLPASAYVRQSQQSCTSGFQAMNLP----- 351
Db
          391 YECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFA 429
Qу
                      : | : ::| : :||:|||
                                                1 1
          352 ----TSSGDLWILGDVFIREYYVVFDRANNYVAMA 382
Db
RESULT 10
REMSK
renin (EC 3.4.23.15) precursor, renal - mouse
N; Alternate names: angiotensin-forming enzyme; angiotensinogenase; renin 1
C; Species: Mus musculus (house mouse)
C;Date: 30-Jun-1987 #sequence revision 30-Jun-1987 #text change 18-Jun-1999
C; Accession: A00989; S07636; A22766; A22058; I57576; A05137; JH0083
R; Holm, I.; Ollo, R.; Panthier, J.J.; Rougeon, F.
EMBO J. 3, 557-562, 1984
A; Title: Evolution of aspartyl proteases by gene duplication: the mouse renin
gene is organized in two homologous clusters of four exons.
A; Reference number: A00989; MUID: 84182525; PMID: 6370686
A; Accession: A00989
A; Molecule type: DNA
A; Residues: 1-402 < HOL>
A; Cross-references: EMBL:X00850
R; Kim, W.S.; Murakami, K.; Nakayama, K.
Nucleic Acids Res. 17, 9480, 1989
A; Title: Nucleotide sequence of a cDNA coding for mouse Ren1 preprorenin.
A; Reference number: S07636; MUID: 90067953; PMID: 2685761
A; Accession: S07636
A; Molecule type: mRNA
A; Residues: 1-402 <KIM>
A; Cross-references: EMBL: X16642; NID: g53930; PIDN: CAA34636.1; PID: g53931
R; Mullins, J.J.; Burt, D.W.; Windass, J.D.; McTurk, P.; George, H.; Brammar,
W.J.
EMBO J. 1, 1461-1466, 1982
A; Title: Molecular cloning of two distinct renin genes from the DBA/2 mouse.
A; Reference number: A90968; MUID: 84207899; PMID: 6327270
A; Accession: A22766
A; Molecule type: mRNA
A; Residues: 269-314, 'D', 316 < MUL>
R; Panthier, J.J.; Dreyfus, M.; Roux, D.T.L.; Rougeon, F.
Proc. Natl. Acad. Sci. U.S.A. 81, 5489-5493, 1984
A; Title: Mouse kidney and submaxillary gland renin genes differ in their 5'
putative regulatory sequences.
A; Reference number: A22058; MUID: 84298161; PMID: 6089205
A; Accession: A22058
A; Molecule type: DNA
A; Residues: 1-30 < PAN>
R; Field, L.J.; Philbrick, W.M.; Howles, P.N.; Dickinson, D.P.; McGowan, R.A.;
Gross, K.W.
Mol. Cell. Biol. 4, 2321-2331, 1984
A; Title: Expression of tissue-specific Ren-1 and Ren-2 genes of mice:
Comparative analysis of 5'-proximal flanking regions.
A; Reference number: I57576; MUID: 85085936; PMID: 6392850
A; Accession: I57576
```

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A; Status: preliminary; translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-31 <RES>
A; Cross-references: GB: K02800; NID: q200689; PIDN: AAA40044.1; PID: q200690
C; Comment: The only known function of renal renin is to release angiotensin I
from angiotensinogen in the plasma, initiating a cascade of reactions that
produces an elevation of blood pressure and increased sodium retention by the
kidney.
C; Comment: Renal renin is synthesized by the juxtaglomerular cells of the kidney
in response to decreased blood pressure and sodium concentration.
C; Genetics:
A; Gene: Ren-1
A; Introns: 31/2; 81/3; 123/1; 162/3; 228/2; 268/2; 316/3; 349/3
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; blood pressure control; glycoprotein;
hydrolase; kidney; plasma
F;1-21/Domain: signal sequence #status predicted <SIG>
F;22-64/Domain: propeptide #status predicted <PRO>
F:65-402/Product: renin #status predicted <MAT>
F; 69, 139, 320/Binding site: carbohydrate (Asn) (covalent) #status predicted
F;102,287/Active site: Asp #status predicted
 Query Match
                        11.9%; Score 320; DB 1; Length 402;
 Best Local Similarity 28.6%; Pred. No. 3.4e-16;
 Matches 126; Conservative 66; Mismatches 181; Indels 68; Gaps
                                                                       21;
          10 LPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPG-PGTPAERHADGLALALE--- 65
Qу
             :11 1 11
                          :1 1:11 1 1
                                           : |
           6 MPLWALLLL----WSPCTFSLPTRTATFERIPLKKMPSVREILEERGVDMTRLSAEWGV 60
Db
          66 ----PA---LASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAV 118
Qy
                    1:
          61 FTKRPSLTNLTSPVVLTNYL----NTQ-----YYGEIGIGTPPQTFKVIFDTGSANLWV 110
Db
         119 AGTPHSY-----IDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTS 172
Qy
               | | | :::: ||:|
                                       111 PSTKCSRLYLACGIHSLYESSDSSSYMENGSDFTIHYGSGRVKGFLSQDSVTV-GGITVT 169
Db
         173 FLVNIATIFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANI-PNVFSMQ 231
Qy
                  | | | |::|:||: |: : ||:::| : |||:
         170 QTFGEVTELPLIPFML--AKFDGVLGMGFP--AQAVGGVTPVFDHILSQGVLKEEVFSVY 225
Db
         232 MCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLD 291
Qy
                         226 Y----NRGSHLLGGEVVLGGSDPQHYQGNFHYVSISKTDSWQITMKGVSVG--SSTLL 277
Db
         292 CREYNADKAIVDSGTTLLRLPQKVFDAVVEAV-ARASLIPEFSDGFWTGSQLACWTNSET 350
Qy
                 | :||:|:: | :::|: |: |:
                                                            : |
         278 CEEGCA--VVVDTGSSFISAPTSSLKLIMOALGAKEKRIEEY-----VVNC---SQV 324
Db
         351 PWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGL-NYECYRFGISPSTNAL-VIG 408
Qy
                            1:: :: : | |
         325 P--TLPDISFDL----GGRAYTLSSTDYVLQYPNRRDKLCTLALHAMDIPPPTGPVWVLG 378
Db
         409 ATVMEGFYVIFDRAQKRVGFA 429
Qу
             | \cdot | : | \cdot | : | \cdot |
                            1:11
         379 ATFIRKFYTEFDRHNNRIGFA 399
Db
```

```
RESULT 11
S66516
oryzasin (EC 3.4.23.-) precursor - rice
N; Alternate names: aspartic proteinase 1
C; Species: Oryza sativa (rice)
C;Date: 28-Oct-1996 #sequence revision 13-Mar-1997 #text change 20-Jun-2000
C; Accession: S66516; S66517
R; Asakura, T.; Watanabe, H.; Abe, K.; Arai, S.
Eur. J. Biochem. 232, 77-83, 1995
A; Title: Rice aspartic proteinase, oryzasin, expressed during seed ripening and
germination, has a gene organization distinct from those of animal and microbial
aspartic proteinases.
A; Reference number: S66516; MUID: 96048031; PMID: 7556174
A:Accession: S66516
A; Molecule type: DNA
A; Residues: 1-509 < ASA>
A;Cross-references: EMBL:D32165; NID:g511665; PIDN:BAA06876.1; PID:g1030715
A; Accession: S66517
A; Molecule type: mRNA
A; Residues: 1-509 <ASZ>
A;Cross-references: EMBL:D32144; NID:g1255684; PIDN:BAA06875.1; PID:g1711289
C; Comment: The pair of saposin repeat homology domains tagged SAP1 and SAP2
represent a cyclical permutation of a single saposin repeat.
A;Introns: 119/3; 140/1; 171/3; 209/2; 265/3; 279/1; 300/3; 338/3; 360/2; 412/3;
452/3; 482/2
C; Superfamily: oryzasin; saposin repeat homology
C; Keywords: aspartic proteinase; hydrolase
F;1-20/Domain: signal sequence #status predicted <SIG>
F;21-68/Domain: propeptide #status predicted <PRO>
F;68-509/Product: aspartic proteinase 1 #status predicted <MAT>
F;316-361/Domain: saposin repeat homology #status atypical <SAP1>
F;370-420/Domain: saposin repeat homology #status atypical <SAP2>
F;103,290/Active site: Asp #status predicted
                         11.7%; Score 313.5; DB 2; Length 509;
  Query Match
  Best Local Similarity 23.0%; Pred. No. 1.5e-15;
 Matches 127; Conservative 75; Mismatches 179; Indels 171;
                                                                         19;
           3 ALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGLAL 62
Qу
                                                :|| |
             ::| || :| || || :|
                                       1 1
           5 SVALVLLAAVLLQALLPASAEEGLVRIALKKRPIDENSRVAARLSG----EEGARRLGL 59
Db
          63 ALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSN----- 115
Qу
                       60 RGANSLGGGGGEGDIVALKNYMNAQ----YFGEIGVGTPPQKFTVIFDTGSSNLWVPSAK 115
Db
         116 --FAVAGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSF 173
Qу
                                            :::||||: |
         116 CYFSIACFFHS----RYKSGOSSTYOKNGKPAAIQYGTGSIAGFFSEDSVTVGD----- 165
Db
         174 LVNIATIFESENFF----LPGI-----KWNGILGLAYATLAKPSSSLETFFDSLVTQANI 224
Qу
                              11:
                                       |::|||| : :: :
                   :::|
         166 ----LVVKDQEFIEATKEPGLTFMVAKFDGILGLGFQEISVGDA------V 206
Db
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225 PNVFSMOMCG-AGLPVAGSGTN-----GGSLVLGGIEPSLYKGDIWYTPIKEEWYYQI 276
Qу
            207 PVWYKMVEQGLVSEPVFSFWFNRHSDEGEGGEIVFGGMDPSHYKGNHTYVPVSQKGYWQF 266
Db
        277 EILKLEIGGOSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPE---- 331
Qу
            267 EMGDVLIGGKTTGF-CA--SGCSAIADSGTSLLAGPTAIITEINEKIGATGVVSQECKTV 323
Db
        332 -----FSDGF---- 336
Qу
                                                         1:1
        324 VSQYGQQILDLLLAETQPSKICSQVGLCTFDGKHGVSAGIKSVVDDEAGESNGLQSGPMC 383
Db
        337 -----FPKISIYLRD 364
QУ
                   [ : | | | : : |
        384 NACEMAVVWMQNQLAQNKTQDLILNYINQLCDKLPSPMGESSVDCGSLASMPEISFTIGA 443
Db
        365 ENSSRSFRITILPQLYIQPMMGAGLNYECY----RFGISPSTNAL-VIGATVMEGFYVIF 419
Qу
           444 K----KFALKPEEYIL-KVGEGAAAQCISGFTAMDIPPPRGPLWILGDVFMGAYHTVF 496
Db
        420 DRAQKRVGFAAS 431
Qу
            | : |||||
        497 DYGKMRVGFAKS 508
Db
RESULT 12
A24608
gastricsin (EC 3.4.23.3) precursor - rat
N; Alternate names: pepsinogen C
N; Contains: pepsin A (EC 3.4.23.1) precursor
C; Species: Rattus norvegicus (Norway rat)
C;Date: 30-Jun-1988 #sequence revision 05-Aug-1994 #text change 18-Jun-1999
C; Accession: A33510; A24608; C22434; A05145; A61298
R; Ishihara, T.; Ichihara, Y.; Hayano, T.; Katsura, I.; Sogawa, K.; Fujii-
Kuriyama, Y.; Takahashi, K.
J. Biol. Chem. 264, 10193-10199, 1989
A; Title: Primary structure and transcriptional regulation of rat pepsinogen C
gene.
A; Reference number: A33510; MUID: 89255508; PMID: 2722863
A; Accession: A33510
A; Molecule type: DNA
A; Residues: 1-392 <ISH>
A; Cross-references: GB:M25985
R; Ichihara, Y.; Soqawa, K.; Morohashi, K.; Fujii-Kuriyama, Y.; Takahashi, K.
Eur. J. Biochem. 161, 7-12, 1986
A; Title: Nucleotide sequence of a nearly full-length cDNA coding for pepsinogen
of rat gastric mucosa.
A; Reference number: A24608; MUID: 87054020; PMID: 3780741
A:Accession: A24608
A; Molecule type: mRNA
A; Residues: 1-392 <ICH>
A; Cross-references: GB: X04644; NID: q56880; PIDN: CAA28305.1; PID: q56881
R; Ichihara, Y.; Sogawa, K.; Takahashi, K.
J. Biochem. 98, 483-492, 1985
A; Title: Isolation of human, swine, and rat prepepsinogens and calf
preprochymosin, and determination of the primary structures of their NH2-
terminal signal sequences.
```

```
A; Reference number: A22434; MUID: 86059312; PMID: 2415509
A; Accession: C22434
A; Molecule type: protein
A; Residues: 1-19, 'X', 21-23, 'X', 25-29 <IC2>
R; Arai, K.M.; Muto, N.; Tani, S.; Akahane, K.
Biochim. Biophys. Acta 788, 256-261, 1984
A; Title: The N-terminal sequence of rat pepsinogen.
A; Reference number: A05145; MUID: 84257697; PMID: 6743670
A; Accession: A05145
A; Molecule type: protein
A; Residues: 17-30, 'Q', 32-102, 'A', 104-108, 'L', 110-112 < ARA>
A; Experimental source: Wistar strain
R; Ichihara, Y.; Sogawa, K.; Takahashi, K.
J. Biochem. 92, 603-606, 1982
A; Title: Rat gastric prepepsinogen: in vitro synthesis and partial amino-
terminal signal sequence.
A; Reference number: A61298; MUID: 83030750; PMID: 6182139
A; Accession: A61298
A; Molecule type: protein
A; Residues: 1, 'XX', 4-6, 'X', 8-9, 'X', 11, 'X', 13-14, 'XXX', 18-
19, 'X', 21, 'X', 23, 'XX', 26, 'X' <IC3>
C; Comment: This enzyme has more restricted specificity than pepsin A. It is the
major form of pepsinogen in rat gastric mucosa.
C; Genetics:
A;Introns: 20/2; 73/3; 113/1; 152/3; 219/2; 259/2; 309/3; 342/3
A; Note: there are at least two very similar genes for gastricsin in rat
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; gastric juice; hydrolase; protein digestion;
stomach
F;1-16/Domain: signal sequence #status experimental <SIG>
F;17-392/Product: pepsinogen #status experimental <MAT>
F;17-62/Domain: activation peptide #status experimental <ACT>
F;94,280/Active site: Asp #status predicted
F;107-112,270-275,314-347/Disulfide bonds: #status predicted
                          11.6%; Score 313; DB 1; Length 392;
  Query Match
  Best Local Similarity 29.5%; Pred. No. 1.1e-15;
 Matches 105; Conservative 56; Mismatches 139; Indels
                                                                     Gaps
                                                                            16;
                                                                56:
           92 YYLEMLIGTPPQKLQILVDTGSSNFAV-----AGTPHSYIDTYFDTERSSTYRSKGF 143
Qу
              1: |: ||||| :| ||||| |
                                                 | | |:
                                                            1: :|||| ::|
           76 YFGEISIGTPPQNFLVLFDTGSSNLWVSSVYCQSEACTTHA----RFNPSKSSTYYTEGQ 131
Db
          144 DVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFESENFFLPG----IKWNGILG 198
Qу
                     || || || || || :|:
                                         1 1
                                                     -1.11
                                                          - 11
          132 TFSLQYGTGSLTGFFGYDTLTV----QSIQVPNQEFGLSEN--EPGTNFVYAQFDGIMG 184
Db
          199 LAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVAGS--GTNGGSLVLGGIEP 256
QУ
                                 :: :
                                          111 1: 1
                           1
          185 LAYPGLS--SGGATTALQGMLGE----GALSQPLFGVYL---GSQQGSNGGQIVFGGVDK 235
Db
          257 SLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVF 316
Qу
              : | | | : | : | | : | | | |
                                       || || || ||
                                                        : |||:||:|| :| :
          236 NLYTGEITWVPVTQELYWQITIDDFLIGDQASGW-CSSQGC-QGIVDTGTSLLVMPAQYL 293
Db
          317 DAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITIL 376
Qу
                                      ::| : |
                                                     1:1
                            1::1
                :::::
```

```
Db
         294 SELLQTIGAQE--GEYGEYF-----VSCDSVSS-----LPTLSFVL-----NGVQFPLS 335
         377 POLY-IOPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qу
                    : 1
                                        ::1 : : | | | |
            1 1 1 1
         336 PSSYIIQEDNFCMVGLESISLTSESGQPLWILGDVFLRSYYAIFDMGNNKVGLATS 391
Db
RESULT 13
A41443
pepsin (EC 3.4.23.-) precursor, embryonic - chicken
C; Species: Gallus gallus (chicken)
C;Date: 05-Jun-1992 #sequence revision 05-Jun-1992 #text change 21-Jul-2000
C; Accession: A41443
R; Hayashi, K.; Aqata, K.; Mochii, M.; Yasuqi, S.; Equchi, G.; Mizuno, T.
J. Biochem. 103, 290-296, 1988
A; Title: Molecular cloning and the nucleotide sequence of cDNA for embryonic
chicken pepsinogen: phylogenetic relationship with prochymosin.
A; Reference number: A41443; MUID: 88227903; PMID: 3131317
A:Accession: A41443
A; Status: preliminary
A; Molecule type: mRNA
A; Residues: 1-383 <HAY>
A;Cross-references: GB:D00215; NID:q2760810; PIDN:BAA00153.1; PID:q222853
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; hydrolase; protein digestion
 Query Match
                       11.5%; Score 310; DB 2; Length 383;
 Best Local Similarity 26.8%; Pred. No. 1.8e-15;
 Matches 106; Conservative 63; Mismatches 136; Indels
                                                                     15;
                                                          90; Gaps
          56 HA--DGLALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGS 113
Qy
                                              | | | : | | | | | | :: | | | |
               Db
          55 HAFPDVLTVVTEPLL-----NTLDM------EYYGTISIGTPPODFTVVFDTGS 97
         114 SNFAVAG----TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGF 169
Qу
            || | :|
                              98 SNLWVPSVSCTSPACOSHOMFNPSOSSTYKSTGONLSIHYGTGDMEGTVGCDTVTVASLM 157
Db
         170 NTSFLVNIATIFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANI-PNVF 228
Qу
            :|: | ::| | || ||: :|::||||| || :|| : : ||::| :: : |:|
         158 DTNOLFGLST-SEPGOFFV-YVKFDGILGLGYPSLA--ADGITPVFDNMVNESLLEQNLF 213
Db
         229 SMOMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSL 288
Qу
            214 SVYLSREPM-----GSMVVFGGIDESYFTGSINWIPVSYQGYWQISMDSIIVNKQEI 265
Db
         289 NLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNS 348
Qу
                   : :||:|:|:|: | : : ||
         266 ACS----SGCQAIIDTGTSLVAGPASDINDIQSAVG------ANQ 300
Db
         349 ETPWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGLNY-----ECY 394
Qу
                            301 NTYGEY-----SVNCSHILAMPDVVF--VIG-GIOYPVPALAYTEQNGQGTCM 345
Db
Qу
         395 RFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFA 429
                : [ : :: | : : | | | | | | | |
         346 SSFQNSSADLWILGDVFIRVYYSIFDRANNRVGLA 380
Db
```

```
RESULT 14
KHHUD
cathepsin D (EC 3.4.23.5) precursor [validated] - human
N; Alternate names: preprocathepsin D
C; Species: Homo sapiens (man)
C;Date: 28-Dec-1987 #sequence_revision 28-Dec-1987 #text_change 15-Sep-2000
C; Accession: A25771; S30749; PC2066; I59236; I57716
R; Faust, P.L.; Kornfeld, S.; Chirgwin, J.M.
Proc. Natl. Acad. Sci. U.S.A. 82, 4910-4914, 1985
A; Title: Cloning and sequence analysis of cDNA for human cathepsin D.
A; Reference number: A25771; MUID: 85270436; PMID: 3927292
A:Accession: A25771
A; Molecule type: mRNA
A; Residues: 1-412 <FAU>
A;Cross-references: EMBL:M11233; NID:q181179; PIDN:AAB59529.1; PID:q181180
R; Westley, B.R.; May, F.E.B.
Nucleic Acids Res. 15, 3773-3786, 1987
A; Title: Oestrogen regulates cathepsin D mRNA levels in oestrogen responsive
human breast cancer cells.
A; Reference number: S30749; MUID: 87231068; PMID: 3588310
A; Accession: S30749
A; Molecule type: mRNA
A; Residues: 1-412 <WES>
A; Cross-references: EMBL: X05344; NID: g29677; PIDN: CAA28955.1; PID: g29678
R; May, F.E.B.; Smith, D.J.; Westley, B.R.
Gene 134, 277-282, 1993
A; Title: The human cathepsin D-encoding gene is transcribed from an estrogen-
regulated and a constitutive start point.
A; Reference number: PC2066; MUID: 94085791; PMID: 8262386
A; Accession: PC2066
A; Molecule type: DNA
A; Residues: 1-23 <MAY>
A;Cross-references: GB:L12980; NID:q291930; PIDN:AAA16314.1; PID:q455429
A; Experimental source: MCF-7 cell
R; Cavailles, V.; Augereau, P.; Rochefort, H.
Proc. Natl. Acad. Sci. U.S.A. 90, 203-207, 1993
A; Title: Cathepsin D gene is controlled by a mixed promoter, and estrogens
stimulate only TATA-dependent transcription in breast cancer cells.
A; Reference number: I59236; MUID: 93126342; PMID: 8419924
A; Accession: I59236
A; Status: translation not shown; translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-22 <CAV1>
A;Cross-references: GB:S52557; NID:g263124; PIDN:AAD13868.1; PID:g4261568
R; Augereau, P.; Miralles, F.; Cavailles, V.; Gaudelet, C.; Parker, M.;
Rochefort, H.
Mol. Endocrinol. 8, 693-703, 1994
A;Title: Characterization of the proximal estrogen-responsive element of human
cathepsin D gene.
A; Reference number: I57716; MUID: 95021301; PMID: 7935485
A; Accession: I57716
A;Status: translation not shown; translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-22 <CAV2>
A;Cross-references: GB:S74689; NID:g786350; PIDN:AAD14156.1; PID:g4261856
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```
R; Baldwin, E.T.; Bhat, T.N.; Gulnik, S.; Erickson, J.W.
submitted to the Brookhaven Protein Data Bank, April 1993
A; Reference number: A51839; PDB:1LYA
A; Contents: annotation; X-ray crystallography, 2.5 angstroms, residues 65-
161;170-241
R; Baldwin, E.T.; Bhat, T.N.; Gulnik, S.; Erickson, J.W.
submitted to the Brookhaven Protein Data Bank, April 1993
A; Reference number: A51840; PDB:1LYB
A; Contents: annotation; X-ray crystallography, 2.5 angstroms, with inhibitor
residues 65-161;170-241
R; Baldwin, E.T.; Bhat, T.N.; Gulnik, S.; Hosur, M.V.; Sowder II, R.C.; Cachau,
R.E.; Collins, J.; Silva, A.M.; Erickson, J.W.
Proc. Natl. Acad. Sci. U.S.A. 90, 6796-6800, 1993
A; Title: Crystal structures of native and inhibited forms of human cathepsin D:
implications for lysosomal targeting and drug design.
A; Reference number: A48229; MUID: 93342076; PMID: 8393577
A; Contents: annotation; X-ray crystallography, 2.5 angstroms
C; Comment: Cathepsin D is a ubiquitous lysosomal proteinase.
C; Comment: In addition to the propeptide, residues 163-168 and 411-412 are
proteolytically removed. Residues 169 and 170 are also partially removed.
C; Comment: The carbohydrate bound to 134-Asn contains a mannose-6-phosphate that
is bound near 267-Lys and the phosphotransferase recognition region.
C; Genetics:
A; Gene: GDB: CTSD
A; Cross-references: GDB:120512; OMIM:116840
A; Map position: 11p15.5-11p15.5
C; Function:
A; Description: limited specificity endopeptidase
A; Pathway: intracellular protein degradation
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; glycoprotein; hydrolase; lysosome; protein
degradation
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F;21-64/Domain: propeptide #status predicted <PRO>
F;65-162,169-410/Product: cathepsin D #status experimental <MAT>
F;267,329-356/Region: phosphotransferase recognition
F;91-160,110-117,286-290,329-366/Disulfide bonds: #status experimental
F;97,295/Active site: Asp #status experimental
F;134,263/Binding site: carbohydrate (Asn) (covalent) #status experimental
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Qy
                      : | :: |
                                                              | :::
            6 LLPLAL--CLLAAP--ASALVRIPLHKFTSIRRTMSEVGGSVEDLIAKGPVSKYSQAVPA 61
Db
           63 ALEPALASPAGAANFLAMVDNLOGDSGRGYYLEMLIGTPPOKLOILVDTGSSNFAVAGTP 122
Qу
                                           1| |: ||||| :: ||||||
                          1::
           62 VTEGPI--PEVLKNYM------DAQYYGEIGIGTPPQCFTVVFDTGSSNLWVPSIH 109
Db
          123 HSYIDT----YFDTERSSTYRSKGFDVTVKYTOGSWTGFVGEDLVTIP--KGFNTSFL 174
Qy
                          :::::||||
                                           : | || :|:: :| |::|
                                     110 CKLLDIACWIHHKYNSDKSSTYVKNGTSFDIHYGSGSLSGYLSQDTVSVPCQSASSASAL 169
Db
          175 --VNIATIFESENFFLPGI----KWNGILGLAYATLAKPSSSLETFFDSLVTQANI-PN 226
Qу
```

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- 1
                            111
                                     1::||||:|| :: :::
                                                             11:1: | : |
         170 GGVKVEROVFGEATKOPGITFIAAKFDGILGMAYPRIS--VNNVLPVFDNLMQQKLVDQN 227
Db
          227 VFSMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQ 286
Qу
                       || |:||| : ||| : | : |:|: : ::|:
Db
         228 IFSFY----LSRDPDAQPGGELMLGGTDSKYYKGSLSYLNVTRKAYWQVHLDQVEV-AS 281
          287 SLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWT 346
Qу
                          :|||:|:|:
                                          : [
                                                              :
          282 GLTL-CKE--GCEAIVDTGTSLMVGP---VDEVRELQKAIGAVPLIQGEY----MIPC-- 329
Db
         347 NSETPWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRF---GISPSTN 403
Qу
                   | | |:: |
                                   ::::: 1: 1 : 11
                                                              330 ---EKVSTLPAITLKL----GGKGYKLS--PEDYTLKVSQAGKTLCLSGFMGMDIPPPSG 380
Db
          404 AL-VIGATVMEGFYVIFDRAQKRVGFA 429
Qy
               1111
          381 PLWILGDVFIGRYYTVFDRDNNRVGFA 407
Db
RESULT 15
KHMSD
cathepsin D (EC 3.4.23.5) precursor - mouse
C; Species: Mus musculus (house mouse)
C;Date: 31-Dec-1991 #sequence_revision 31-Dec-1991 #text change 18-Jun-1999
C; Accession: I48278; S14704; S12587
R; Hetman, M.; Perschl, A.; Saftig, P.; Von Figura, K.; Peters, C.
DNA Cell Biol. 13, 419-427, 1994
A; Title: Mouse cathepsin D gene: molecular organization, characterization of the
promoter, and chromosomal localization.
A; Reference number: I48278; MUID: 94280622; PMID: 8011168
A; Accession: I48278
A; Status: preliminary; translated from GB/EMBL/DDBJ
A; Molecule type: DNA
A; Residues: 1-410 < RES>
A; Cross-references: EMBL: X68378; NID: g50302; PIDN: CAA48453.1; PID: g817945
R; Diedrich, J.F.; Staskus, K.A.; Retzel, E.F.; Haase, A.T.
Nucleic Acids Res. 18, 7184, 1990
A; Title: Nucleotide sequence of a cDNA encoding mouse cathepsin D.
A; Reference number: S14704; MUID: 91088345; PMID: 2263503
A; Accession: S14704
A; Molecule type: mRNA
A; Residues: 1-410 <DIE>
A; Cross-references: EMBL: X53337; NID: g50300; PIDN: CAA37423.1; PID: g50301
R; Grusby, M.J.; Mitchell, S.C.; Glimcher, L.H.
Nucleic Acids Res. 18, 4008, 1990
A; Title: Molecular cloning of mouse cathepsin D.
A; Reference number: S12587; MUID: 90326544; PMID: 2374732
A; Accession: S12587
A; Molecule type: mRNA
A; Residues: 1-410 <GRU>
A;Cross-references: EMBL:X52886; NID:g50298; PIDN:CAA37067.1; PID:g50299
C: Genetics:
A; Introns: 23/2; 76/3; 118/1; 157/3; 233/2; 274/2; 322/3; 355/3
C; Function:
A; Description: limited specificity endopeptidase
A; Pathway: intracellular protein degradation
```

```
C; Superfamily: pepsin
C; Keywords: aspartic proteinase; glycoprotein; hydrolase; lysosome; protein
F;1-20/Domain: signal sequence #status predicted <SIG>
F;21-64/Domain: propeptide #status predicted <PRO>
F;65-410/Product: cathepsin D, single-chain form #status predicted <MAT>
F;91-160,110-117,284-288,327-364/Disulfide bonds: #status predicted
F:97,293/Active site: Asp #status predicted
F;134,261/Binding site: carbohydrate (Asn) (covalent) #status predicted
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 Query Match
 Best Local Similarity 27.5%; Pred. No. 3.6e-15;
 Matches 103; Conservative 64; Mismatches 123; Indels 85; Gaps
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Qу
            79 YYGDIGIGTPPQCFTVVFDTGSSNLWVPSIHCKILDIACWVHHKYNSDKSSTYVKNGTSF 138
Db
        146 TVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIAT---IFESENFFLPGI----KWNGIL 197
Qу
             139 DIHYGSGSLSGYLSQDTVSVPCKSDQSKARGIKVEKQIF-GEATKQPGIVFVAAKFDGIL 197
Db
        198 GLAYATLAKPSSSLETFFDSLVTQANI-PNVFSMQMCGAGLPVAGSGTNGGSLVLGGIEP 256
Qy
            198 GMGYPHIS--VNNVLPVFDNLMQQKLVDKNIFSFY----LNRDPEGQPGGELMLGGTDS 250
Db
        257 SLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVF 316
Qу
              1 |:: | : : |:|: : : ||:| : | | |:
                                               : | | | | : | | : | | : |
        251 KYYHGELSYLNVTRKAYWQVHMDQLEVGNE-LTL-CK--GGCEAIVDTGTSLLVGPVEEV 306
Db
        317 DAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITIL 376
Qγ
                                                   : |
             :: |: ||
        307 KELOKAIGAVPLI------QGEYMIPCEKVSSL 333
Db
        377 PQLYIQPMMGAGLNYEC----YRFGIS------PSTNALVIGATVMEG 414
Qy
            334 PTVYLK--LG-GKNYELHPDKYILKVSQGGKTICLSGFMGMDIPPPSGPLWILGDVFIGS 390
Db
        415 FYVIFDRAQKRVGFA 429
Qу
            :|:||| |||
        391 YYTVFDRDNNRVGFA 405
Db
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Search completed: March 4, 2004, 15:40:59 Job time: 29.1043 secs

# GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

March 4, 2004, 15:39:01; Search time 57.8617 Seconds Run on:

(without alignments)

1890.324 Million cell updates/sec

US-09-668-314C-2 Title:

Perfect score: 2687

1 MGALARALLLPLLAQWLLRA......RPRDPEVVNDESSLVRHRWK 518 Sequence:

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

809742 segs, 211153259 residues Searched:

809742 Total number of hits satisfying chosen parameters:

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database : Published Applications AA:\*

/cgn2\_6/ptodata/2/pubpaa/US07\_PUBCOMB.pep:\*

/cgn2 6/ptodata/2/pubpaa/PCT NEW PUB.pep:\*

/cgn2 6/ptodata/2/pubpaa/US06 NEW\_PUB.pep:\*

/cgn2 6/ptodata/2/pubpaa/US06 PUBCOMB.pep:\*

/cgn2 6/ptodata/2/pubpaa/US07 NEW PUB.pep:\*

/cgn2 6/ptodata/2/pubpaa/PCTUS PUBCOMB.pep:\*

/cgn2 6/ptodata/2/pubpaa/US08\_NEW\_PUB.pep:\*

7: /cgn2 6/ptodata/2/pubpaa/US08 PUBCOMB.pep:\*

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10: /cgn2\_6/ptodata/2/pubpaa/US09B\_PUBCOMB.pep:\*

11: /cgn2 6/ptodata/2/pubpaa/US09C PUBCOMB.pep:\*

12: /cgn2\_6/ptodata/2/pubpaa/US09 NEW PUB.pep:\*

13: /cgn2 6/ptodata/2/pubpaa/US10A PUBCOMB.pep:\*

14: /cgn2 6/ptodata/2/pubpaa/US10B PUBCOMB.pep:\*

15: /cgn2 6/ptodata/2/pubpaa/US10C PUBCOMB.pep:\*

16: /cgn2 6/ptodata/2/pubpaa/US10 NEW PUB.pep:\*

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Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

કૃ

Result Query

> No. Score Match Length DB ID

Description

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# ALIGNMENTS

# RESULT 1 US-09-794-927-2

- ; Sequence 2, Application US/09794927 ; Patent No. US20010016324A1
- ; GENERAL INFORMATION:
- ; APPLICANT: Gurney, Mark E.

```
APPLICANT: Bienkowski, Michael J.
            Heinrikson, Robert L.
  APPLICANT:
            Parodi, Luis A.
  APPLICANT:
            Yan, Rigiang
  APPLICANT:
  TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR,
AND
  TITLE OF INVENTION: USES
  TITLE OF INVENTION:
                   THEREFOR
  FILE REFERENCE: 28341/6280FG
  CURRENT APPLICATION NUMBER: US/09/794,927
  CURRENT FILING DATE: 2001-02-27
  PRIOR APPLICATION NUMBER: 09/416,901
  PRIOR FILING DATE: 1999-10-13
  PRIOR APPLICATION NUMBER: 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: 60/101,594
  PRIOR FILING DATE: 1998-09-24
  NUMBER OF SEQ ID NOS: 73
  SOFTWARE: PatentIn Ver. 2.0
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   ORGANISM: Homo sapiens
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                                             Length 518;
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; Patent No. US20010018208A1
; GENERAL INFORMATION:
  APPLICANT: Gurney, Mark E.
  APPLICANT: Bienkowski, Michael J.
  APPLICANT: Heinrikson, Robert L.
             Parodi, Luis A.
  APPLICANT:
  APPLICANT: Yan, Rigiang
  TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR,
AND
  TITLE OF INVENTION: USES
  TITLE OF INVENTION: THEREFOR
  FILE REFERENCE: 28341/6280DE
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  CURRENT FILING DATE: 2001-02-28
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  PRIOR FILING DATE: 1999-10-13
  PRIOR APPLICATION NUMBER: 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: 60/101,594
  PRIOR FILING DATE: 1998-09-24
  NUMBER OF SEQ ID NOS: 73
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   LENGTH: 518
   TYPE: PRT
   ORGANISM: Homo sapiens
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  Query Match
                       100.0%; Pred. No. 2.8e-240;
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; APPLIC							
; APPLIC ; APPLIC		·					
		Parodi, Luis A.					
; APPLIC	CANT:	Yan, Riqiang					
	OF I	NVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THERE	FOR,				
AND	OF T	INVENTION: USES					
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	CURRENT FILING DATE: 2001-02-27						
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		ING DATE: 1999-09-23					
: PRIOR	APPT	TCATION NUMBER: PCT/US99/20881					

; PRIOR APPLICATION NUMBER: PCT/US99/20881

; PRIOR FILING DATE: 1999-09-23

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  PRIOR FILING DATE: 1998-09-24
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   ORGANISM: Homo sapiens
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                                          Length 518;
 Query Match
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                   100.0%; Pred. No. 2.8e-240;
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- ; Patent No. US20020037315A1
- ; GENERAL INFORMATION:
- ; APPLICANT: Gurney, Mark E.

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Bienkowski, Michael J.
  APPLICANT:
            Heinrikson, Robert L.
  APPLICANT:
            Parodi, Luis A.
  APPLICANT:
            Yan, Riqiang
  APPLICANT:
  TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR,
AND
  TITLE OF INVENTION:
                   USES
  TITLE OF INVENTION:
                   THEREFOR
  FILE REFERENCE: 28341/6280JL
  CURRENT APPLICATION NUMBER: US/09/794,748
  CURRENT FILING DATE: 2001-02-27
  PRIOR APPLICATION NUMBER: 09/416,901
  PRIOR FILING DATE: 1999-10-13
  PRIOR APPLICATION NUMBER: 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: 60/101,594
  PRIOR FILING DATE: 1998-09-24
  NUMBER OF SEQ ID NOS: 73
  SOFTWARE: PatentIn Ver. 2.0
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   LENGTH: 518
   TYPE: PRT
   ORGANISM: Homo sapiens
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; Patent No. US20020064819A1
; GENERAL INFORMATION:
; APPLICANT: Gurney, Mark E.
 APPLICANT: Bienkowski, Michael J.
  APPLICANT: Heinrikson, Robert L.
  APPLICANT: Parodi, Luis A.
  APPLICANT: Yan, Rigiang
  TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR,
AND USES
  TITLE OF INVENTION: THEREFOR
  FILE REFERENCE: 28341/6280HI
  CURRENT APPLICATION NUMBER: US/09/794,925
  CURRENT FILING DATE: 2001-02-27
  PRIOR APPLICATION NUMBER: 09/416,901
  PRIOR FILING DATE: 1999-10-13
  PRIOR APPLICATION NUMBER: 60/155,493
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: 09/404,133
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: PCT/US99/20881
  PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: 60/101,594
  PRIOR FILING DATE: 1998-09-24
  NUMBER OF SEQ ID NOS: 73
  SOFTWARE: PatentIn Ver. 2.0
; SEQ ID NO 2
   LENGTH: 518
   TYPE: PRT
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US-09-794-925-2
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; Patent No. US20020068278A1
; GENERAL INFORMATION:
  APPLICANT: Giese, Klaus
  APPLICANT: Xin, Hong
  TITLE OF INVENTION: METASTATIC BREAST AND COLON CANCER REGULATED GENES
  FILE REFERENCE: 1451.100 / 210030.447
  CURRENT APPLICATION NUMBER: US/09/215,450
  CURRENT FILING DATE: 1998-12-17
  NUMBER OF SEQ ID NOS: 27
  SOFTWARE: FastSEQ for Windows Version 3.0
 SEQ ID NO 19
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US-09-215-450-19
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 Query Match
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 Best Local Similarity
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- ; Sequence 2, Application US/09681442
- ; Patent No. US20020081634A1
- ; GENERAL INFORMATION:
- ; APPLICANT: Gurney, Mark E.
- ; APPLICANT: Bienkowski, Michael J.
- ; APPLICANT: Heinrikson, Robert L.
- APPLICANT: Parodi, Luis A.
- APPLICANT: Yan, Riqianq
- TITLE OF INVENTION: ALZHEIMER'S DISEASE SECRETASE, APP SUBSTRATES THEREFOR, AND USES
- TITLE OF INVENTION: THEREFOR
- ; FILE REFERENCE: 28341/6280FG
- CURRENT APPLICATION NUMBER: US/09/681,442
- CURRENT FILING DATE: 2001-04-05
- PRIOR APPLICATION NUMBER: 09/416,901
- PRIOR FILING DATE: 1999-10-13
- PRIOR APPLICATION NUMBER: 60/155,493
- PRIOR FILING DATE: 1999-09-23
- PRIOR APPLICATION NUMBER: 09/404,133
- ; PRIOR FILING DATE: 1999-09-23
- ; PRIOR APPLICATION NUMBER: PCT/US99/20881

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PRIOR FILING DATE: 1999-09-23
  PRIOR APPLICATION NUMBER: 60/101,594
  PRIOR FILING DATE: 1998-09-24
  NUMBER OF SEQ ID NOS: 73
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   ORGANISM: Homo sapiens
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### RESULT 8

US-09-978-295A-196

- ; Sequence 196, Application US/09978295A
- ; Patent No. US20020156006A1
- ; GENERAL INFORMATION:

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APPLICANT: Ashkenazi, Avi
APPLICANT: Baker Kevin P.
            Botstein, David
APPLICANT:
            Desnoyers, Luc
APPLICANT:
            Eaton, Dan
APPLICANT:
APPLICANT: Ferrara, Napoleon
APPLICANT: Filvaroff, Ellen
APPLICANT: Fong, Sherman
APPLICANT: Gao, Wei-Qiang
APPLICANT: Gerber, Hanspeter
APPLICANT: Gerritsen, Mary E.
APPLICANT: Goddard, Audrey
APPLICANT: Godowski, Paul J.
APPLICANT: Grimaldi, J. Christopher
APPLICANT: Gurney, Austin L.
APPLICANT: Hillan, Kenneth J
APPLICANT: Kljavin, Ivar J.
APPLICANT: Kuo, Sophia S.
APPLICANT:
            Napier, Mary A.
            Pan, James;
APPLICANT:
           Paoni, Nicholas F.
APPLICANT:
APPLICANT: Roy, Margaret Ann
APPLICANT: Shelton, David L.
APPLICANT: Stewart, Timothy A.
            Tumas, Daniel
APPLICANT:
            Williams, P. Mickey
APPLICANT:
APPLICANT: Wood, William I.
TITLE OF INVENTION: Secreted and Transmembrane Polypeptides and Nucleic
TITLE OF INVENTION: Acids Encoding the Same
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CURRENT FILING DATE: 2001-10-15
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 PRIOR FILING DATE: 2001-07-30
 PRIOR APPLICATION NUMBER: 60/062250
 PRIOR FILING DATE: 1997-10-17
 PRIOR APPLICATION NUMBER: 60/064249
 PRIOR FILING DATE: 1997-11-03
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PRIOR APPLICATION NUMBER: 60/084414

PRIOR APPLICATION NUMBER: 60/084441 PRIOR FILING DATE: 1998-05-06

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  PRIOR FILING DATE: 1998-05-15
  PRIOR APPLICATION NUMBER: 60/085697
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; Patent No. US20020159991A1
; GENERAL INFORMATION:
  APPLICANT: Cordell, Barbara
  APPLICANT: Schimmoller, Frauke
            Liu, Yu-Wang
  APPLICANT:
  APPLICANT: Quon, Diana Hom
  TITLE OF INVENTION: Modulation of A Levels by
  TITLE OF INVENTION: Secretase BACE2
  FILE REFERENCE: SCIOS.022A
  CURRENT APPLICATION NUMBER: US/09/886,143
  CURRENT FILING DATE: 2001-06-20
  PRIOR APPLICATION NUMBER: 60/215,729
  PRIOR FILING DATE: 2000-06-28
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RESULT 10
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; Sequence 196, Application US/09978697
; Patent No. US20020169284A1
; GENERAL INFORMATION:
  APPLICANT: Ashkenazi, Avi
 APPLICANT: Baker Kevin P.
  APPLICANT: Botstein, David
  APPLICANT: Desnoyers, Luc
; APPLICANT: Eaton, Dan
; APPLICANT: Ferrara, Napoleon
; APPLICANT: Filvaroff, Ellen
  APPLICANT: Fong, Sherman
; APPLICANT: Gao, Wei-Qiang
  APPLICANT: Gerber, Hanspeter
  APPLICANT: Gerritsen, Mary E.
  APPLICANT: Goddard, Audrey
  APPLICANT: Godowski, Paul J.
  APPLICANT: Grimaldi, J. Christopher
  APPLICANT: Gurney, Austin L.
  APPLICANT: Hillan, Kenneth J
  APPLICANT: Kljavin, Ivar J.
  APPLICANT: Kuo, Sophia S.
  APPLICANT: Napier, Mary A.
  APPLICANT: Pan, James;
   APPLICANT: Paoni, Nicholas F.
  APPLICANT: Roy, Margaret Ann
   APPLICANT: Shelton, David L.
  APPLICANT: Stewart, Timothy A.
  APPLICANT: Tumas, Daniel
  APPLICANT: Williams, P. Mickey
  APPLICANT: Wood, William I.
   TITLE OF INVENTION: Secreted and Transmembrane Polypeptides and Nucleic
   TITLE OF INVENTION: Acids Encoding the Same
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  CURRENT FILING DATE: 2001-10-16
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Qу	121	TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI	180
Db	121	TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI	180
Qу	181	FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA	240
Db	181	FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA	240
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Qу	301	IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI	360
Db	301	IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI	360
Qу	361	YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD	420
Db	361	YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD	420
Qу	421	RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG	480
Db	421	RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG	480
Qу	481	AILLVLIVLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518	
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## RESULT 11

US-09-978-192A-196

- ; Sequence 196, Application US/09978192A
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- ; GENERAL INFORMATION:
- ; APPLICANT: Ashkenazi, Avi
- ; APPLICANT: Baker Kevin P.
- ; APPLICANT: Botstein, David
- ; APPLICANT: Desnoyers, Luc
- ; APPLICANT: Eaton, Dan
- ; APPLICANT: Ferrara, Napoleon
- ; APPLICANT: Filvaroff, Ellen
- ; APPLICANT: Fong, Sherman
- ; APPLICANT: Gao, Wei-Qiang
- ; APPLICANT: Gerber, Hanspeter
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 APPLICANT: Tumas, Daniel
 APPLICANT: Williams, P. Mickey
 APPLICANT: Wood, William I.
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 TITLE OF INVENTION: Acids Encoding the Same
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#### RESULT 12

US-09-999-832A-196

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  APPLICANT: Shelton, David L.
; APPLICANT: Stewart, Timothy A.
; APPLICANT: Tumas, Daniel
   APPLICANT: Williams, P. Mickey
   APPLICANT: Wood, William I.
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   TITLE OF INVENTION: Acids Encoding the Same
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  PRIOR FILING DATE: 1998-05-13
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  PRIOR FILING DATE: 1998-05-15
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  PRIOR FILING DATE: 1998-05-15
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  PRIOR FILING DATE: 1998-05-15
  PRIOR APPLICATION NUMBER: 60/085580
  PRIOR FILING DATE: 1998-05-15
  PRIOR APPLICATION NUMBER: 60/085573
  PRIOR FILING DATE: 1998-05-15
  PRIOR APPLICATION NUMBER: 60/085704
  PRIOR FILING DATE: 1998-05-15
  PRIOR APPLICATION NUMBER: 60/085697
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                             Score 2687; DB 9; Length 518;
 Query Match
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                             Pred. No. 2.8e-240;
                     100.0%;
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                           0; Mismatches
                                                      0;
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                                                         Gaps
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Qу
           61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
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QУ
           121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qy
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           241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
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Qу
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US-09-978-189-196
; Sequence 196, Application US/09978189
; Publication No. US20030004102A1
; GENERAL INFORMATION:
  APPLICANT: Ashkenazi, Avi
  APPLICANT: Baker Kevin P.
  APPLICANT: Botstein, David
  APPLICANT: Desnoyers, Luc
 APPLICANT: Eaton, Dan
 APPLICANT: Ferrara, Napoleon
 APPLICANT: Filvaroff, Ellen
 APPLICANT: Fong, Sherman
 APPLICANT: Gao, Wei-Qiang
 APPLICANT: Gerber, Hanspeter
  APPLICANT: Gerritsen, Mary E.
  APPLICANT: Goddard, Audrey
  APPLICANT: Godowski, Paul J.
  APPLICANT: Grimaldi, J. Christopher
 APPLICANT: Gurney, Austin L.
; APPLICANT: Hillan, Kenneth J
 APPLICANT: Kljavin, Ivar J.
 APPLICANT: Kuo, Sophia S.
 APPLICANT: Napier, Mary A.
 APPLICANT: Pan, James;
 APPLICANT: Paoni, Nicholas F.
 APPLICANT: Roy, Margaret Ann
  APPLICANT: Shelton, David L.
  APPLICANT: Stewart, Timothy A.
  APPLICANT:
             Tumas, Daniel
 APPLICANT: Williams, P. Mickey
 APPLICANT: Wood, William I.
  TITLE OF INVENTION: Secreted and Transmembrane Polypeptides and Nucleic
 TITLE OF INVENTION: Acids Encoding the Same
 FILE REFERENCE: P2630P1C7
  CURRENT APPLICATION NUMBER: US/09/978,189
  CURRENT FILING DATE: 2001-10-15
  PRIOR APPLICATION NUMBER: 09/918585
  PRIOR FILING DATE: 2001-07-30
  PRIOR APPLICATION NUMBER: 60/062250
; PRIOR FILING DATE: 1997-10-17
; PRIOR APPLICATION NUMBER: 60/064249
; PRIOR FILING DATE: 1997-11-03
; PRIOR APPLICATION NUMBER: 60/065311
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PRIOR FILING DATE: 1997-11-21

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- ; PRIOR FILING DATE: 1998-03-10
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- ; PRIOR APPLICATION NUMBER: 60/081049

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  PRIOR APPLICATION NUMBER: 60/085582
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  PRIOR APPLICATION NUMBER: 60/085700
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  PRIOR APPLICATION NUMBER: 60/085689
  PRIOR FILING DATE: 1998-05-15
  PRIOR APPLICATION NUMBER: 60/085579
  PRIOR FILING DATE: 1998-05-15
  PRIOR APPLICATION NUMBER: 60/085580
  PRIOR FILING DATE: 1998-05-15
  PRIOR APPLICATION NUMBER: 60/085573
  PRIOR FILING DATE: 1998-05-15
  PRIOR APPLICATION NUMBER: 60/085704
  PRIOR FILING DATE: 1998-05-15
  PRIOR APPLICATION NUMBER: 60/085697
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 Best Local Similarity
 Matches 518; Conservative
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Qу
             1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
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US-09-978-608A-196

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- ; Publication No. US20030045462A1
- ; GENERAL INFORMATION:
- ; APPLICANT: Ashkenazi, Avi
- ; APPLICANT: Baker Kevin P.
- ; APPLICANT: Botstein, David
- ; APPLICANT: Desnoyers, Luc
- ; APPLICANT: Eaton, Dan
- ; APPLICANT: Ferrara, Napoleon
- : APPLICANT: Filvaroff, Ellen
- ; APPLICANT: Fong, Sherman
- ; APPLICANT: Gao, Wei-Qiang
- ; APPLICANT: Gerber, Hanspeter
- APPLICANT: Gerritsen, Mary E.
- ; APPLICANT: Goddard, Audrey
- ; APPLICANT: Godowski, Paul J.
- ; APPLICANT: Grimaldi, J. Christopher
- ; APPLICANT: Gurney, Austin L.
- ; APPLICANT: Hillan, Kenneth J
- ; APPLICANT: Kljavin, Ivar J.
- ; APPLICANT: Kuo, Sophia S.
- ; APPLICANT: Napier, Mary A.
- ; APPLICANT: Pan, James;
- ; APPLICANT: Paoni, Nicholas F.
- ; APPLICANT: Roy, Margaret Ann
- ; APPLICANT: Shelton, David L.
- APPLICANT: Stewart, Timothy A.
- ; APPLICANT: Tumas, Daniel
- ; APPLICANT: Williams, P. Mickey

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APPLICANT: Wood, William I.
  TITLE OF INVENTION: Secreted and Transmembrane Polypeptides and Nucleic
  TITLE OF INVENTION: Acids Encoding the Same
  FILE REFERENCE: P2630P1C22
  CURRENT APPLICATION NUMBER: US/09/978,608A
  CURRENT FILING DATE: 2001-10-16
  NUMBER OF SEQ ID NOS: 624
 Prior Application removed - See File Wrapper or Palm
 SEQ ID NO 196
   LENGTH: 518
   TYPE: PRT
   ORGANISM: Homo sapien
US-09-978-608A-196
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 Matches 518; Conservative
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           1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
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; Sequence 196, Application US/09978585A
; Publication No. US20030049633A1
; GENERAL INFORMATION:
  APPLICANT: Ashkenazi, Avi
  APPLICANT: Baker Kevin P.
  APPLICANT: Botstein, David
  APPLICANT: Desnoyers, Luc
  APPLICANT: Eaton, Dan
  APPLICANT: Ferrara, Napoleon
  APPLICANT: Filvaroff, Ellen
  APPLICANT: Fong, Sherman APPLICANT: Gao, Wei-Qiang
  APPLICANT: Gerber, Hanspeter
  APPLICANT: Gerritsen, Mary E.
  APPLICANT: Goddard, Audrey
  APPLICANT: Godowski, Paul J.
  APPLICANT: Grimaldi, J. Christopher
 APPLICANT: Gurney, Austin L.
  APPLICANT: Hillan, Kenneth J
  APPLICANT: Kljavin, Ivar J.
  APPLICANT: Kuo, Sophia S.
  APPLICANT: Napier, Mary A.
  APPLICANT: Pan, James;
  APPLICANT: Paoni, Nicholas F.
  APPLICANT: Roy, Margaret Ann
  APPLICANT: Shelton, David L.
  APPLICANT: Stewart, Timothy A.
  APPLICANT:
             Tumas, Daniel
  APPLICANT:
             Williams, P. Mickey
  APPLICANT:
             Wood, William I.
  TITLE OF INVENTION: Secreted and Transmembrane Polypeptides and Nucleic
  TITLE OF INVENTION: Acids Encoding the Same
  FILE REFERENCE: P2630P1C15
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  CURRENT FILING DATE: 2001-10-16
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; Prior Application removed - See File Wrapper or Palm
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US-09-978-585A-196
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 Best Local Similarity
                       100.0%; Pred. No. 2.8e-240;
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Search completed: March 4, 2004, 15:57:36

Job time: 59.8617 secs

# GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

March 4, 2004, 15:28:35; Search time 74.3936 Seconds Run on:

(without alignments)

2196.942 Million cell updates/sec

Title: US-09-668-314C-2

Perfect score: 2687

Sequence: 1 MGALARALLLPLLAQWLLRA......RPRDPEVVNDESSLVRHRWK 518

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

1017041 seqs, 315518202 residues Searched:

Total number of hits satisfying chosen parameters: 1017041

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database : SPTREMBL 25:\*

1: sp\_archea:\*
2: sp\_bacteria:\*

3: sp\_fungi:\*

4: sp\_human:\*

5: sp invertebrate:\*

6: sp mammal:\*

7: sp mhc:\*

8: sp organelle:\*

9: sp phage:\*

10: sp\_plant:\*

11: sp\_rodent:\*
12: sp\_virus:\*

13: sp\_vertebrate:\*

14: sp\_unclassified:\*

15: sp rvirus:\*

16: sp bacteriap:\*

17: sp archeap:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## SUMMARIES

કૃ

Result Query

No. Score Match Length DB ID

Description

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2	2399	89.3	514	11	Q8C793	Q8c793 mus musculu
3	2395	89.1	514	11	Q9JL18	Q9jl18 mus musculu
4	2375	88.4	468	4	Q9NZL2	Q9nzl2 homo sapien
5	2293	85.3	439	4	Q9H2V8	Q9h2v8 homo sapien
6	2184	81.3	423	4	Q8N2D4	Q8n2d4 homo sapien
7	1966	73.2	396	4	Q9NZL1	Q9nzl1 homo sapien
8	1693	63.0	500	13	Q7T0Y2	Q7t0y2 xenopus lae
9	1246	46.4	255	11	Q9R1P7	Q9r1p7 mus musculu
10	1186	44.1	501	11	Q8C7R1	Q8c7r1 mus musculu
11	1183.5	44.0	532	4	Q9ULS1	Q9uls1 homo sapien
12	1183	44.0	501	11	Q8BQY4	Q8bqy4 mus musculu
13	1172.5	43.6	501	4	Q8IYC8	Q8iyc8 homo sapien
$\frac{13}{14}$	1049	39.0	467	11	Q8C4F4	Q8c4f4 mus musculu
15	653	24.3	267	11	Q0C414 Q9CUU5	Q0C414 mus musculu Q9cuu5 mus musculu
					-	~
16	461	17.2	213	4	Q9P0D2	Q9p0d2 homo sapien
17	386	14.4	244	5	Q8WQY9	Q8wqy9 aphrocallis
18	367.5	13.7	383	13	Q9DEC3	Q9dec3 xenopus lae
19	361.5	13.5	389	6	Q9GMY4	Q9gmy4 sorex ungui
20	355.5	13.2	384	13	Q91322	Q91322 rana catesb
21	355	13.2	389	13	Q9W643	Q9w643 gallus gall
22	355	13.2	389	13	Q9PWK1	Q9pwk1 gallus gall
23	351.5	13.1	389	6	Q9GMY3	Q9gmy3 rhinolophus
24	351.5	13.1	389	6	Q9GMY5	Q9gmy5 suncus muri
25	335.5	12.5	388	6	Q9GMY2	Q9gmy2 oryctolagus
26	334.5	12.4	391	5	Q9VKP6	Q9vkp6 drosophila
27	326	12.1	399	13	093458	O93458 podarcis si
28	324.5	12.1	372	5	Q9VLK3	Q9vlk3 drosophila
29	324.5	12.1	383	13	Q9DE45	Q9de45 salvelinus
30	322.5	12.0	390	6	Q8SQ41	Q8sq41 canis famil
31	322.5	12.0	397	13	Q9W6D4	Q9w6d4 hynobius le
32	320	11.9	385	13	Q9DEC4	Q9dec4 rana catesb
33	319.5	11.9	387	13	Q9DDV5	Q9ddv5 salvelinus
34	319	11.9	419	5	Q95VA2	Q95va2 clonorchis
35	318.5	11.9	396	13	093428	093428 chionodraco
36	318	11.8	397	13	Q800A0	Q800a0 rana catesb
37	316	11.8	378	13	Q9PUR9	Q9pur9 pseudopleur
38	316	11.8	392	11	Q9D7R7	Q9d7r7 mus musculu
39	313	11.6	383	5	076856	076856 dictyosteli
40	312.5	11.6	354	5	Q9GYX7	Q9qyx7 boophilus m
	305	11.4			_	Q9dec2 xenopus lae
41			384	13 13	Q9DEC2	P87370 oncorhynchu
42	302	11.2	398		P87370	Q91x66 mus musculu
43	301.5	11.2	401	11	Q91X66	
44	300.5	11.2	386	6	Q9BGU5	Q9bgu5 bos taurus
45	295.5	11.0	390	6	Q9GK10	Q9gk10 camelus dro

# ALIGNMENTS

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RESULT 1
Q8C5E9

ID Q8C5E9 PRELIMINARY; PRT; 514 AA.

AC Q8C5E9;

DT 01-MAR-2003 (TrEMBLrel. 23, Created)

DT 01-MAR-2003 (TrEMBLrel. 23, Last sequence update)

DT 01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
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Beta-site APP-cleaving enzyme 2.
    BACE2.
GN
    Mus musculus (Mouse).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OC
    NCBI TaxID=10090;
OX
    [1]
RN
RP
    SEQUENCE FROM N.A.
    STRAIN=C57BL/6J; TISSUE=Testis;
RC
    MEDLINE=22354683; PubMed=12466851;
RX
    The FANTOM Consortium,
RA
    the RIKEN Genome Exploration Research Group Phase I & II Team;
RA
    "Analysis of the mouse transcriptome based on functional annotation of
RТ
RТ
    60,770 full-length cDNAs.";
    Nature 420:563-573(2002).
RL
    EMBL; AK078770; BAC37384.1; -.
DR
DR
    MGD; MGI:1860440; Bace2.
    GO; GO:0004194; F:pepsin A activity; IEA.
DR
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR001461; Peptidase A1.
DR
DR
    InterPro; IPR009007; Pept A acid.
    Pfam; PF00026; asp; 1.
DR
DR
    PRINTS; PR00792; PEPSIN.
DR
    PROSITE; PS00141; ASP PROTEASE; 2.
SQ
    SEQUENCE
             514 AA; 55811 MW; CBB9237BB68A0B2E CRC64;
 Query Match
                      89.5%; Score 2405; DB 11; Length 514;
 Best Local Similarity
                      88.8%; Pred. No. 3.5e-179;
 Matches 460; Conservative 20; Mismatches
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Db
         61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qу
                       61 ALALEPVRAT----ANFLAMVDNLOGDSGRGYYLEMLIGTPPOKVOILVDTGSSNFAVAG 116
Db
        121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
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            Db
        117 APHSYIDTYFDSESSSTYHSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNSSFLVNIATI 176
        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qу
           177 FESENFFLPGIKWNGILGLAYAALAKPSSSLETFFDSLVAQAKIPDIFSMQMCGAGLPVA 236
Db
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qу
           237 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQNLNLDCREYNADKA 296
Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qy
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Qу
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357 YLRDENASRSFRITILPQLYIQPMMGAGFNYECYRFGISSSTNALVIGATVMEGFYVVFD 416
Db
         421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Qу
             417 RAQRRVGFAVSPCAEIEGTTVSEISGPFSTEDIASNCVPAQALNEPILWIVSYALMSVCG 476
Db
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Qу
             111111:1111 |: 111111111111111111
         477 AILLVLILLLLPLHCRHAPRDPEVVNDESSLVRHRWK 514
Db
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08C793
                                       514 AA.
                                PRT;
                PRELIMINARY;
    08C793
ID
    08C793;
AC.
    01-MAR-2003 (TrEMBLrel. 23, Created)
DT
     01-MAR-2003 (TrEMBLrel. 23, Last sequence update)
DT
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DT
    Beta-site APP-cleaving enzyme 2.
DE
GN
    BACE2.
    Mus musculus (Mouse).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OC
OX
    NCBI TaxID=10090;
RN
     [1]
     SEQUENCE FROM N.A.
RP
     STRAIN=C57BL/6J; TISSUE=Heart;
RC
     MEDLINE=22354683; PubMed=12466851;
RX
     The FANTOM Consortium,
RA
     the RIKEN Genome Exploration Research Group Phase I & II Team;
RA
     "Analysis of the mouse transcriptome based on functional annotation of
RT
     60,770 full-length cDNAs.";
RT
     Nature 420:563-573(2002).
RL
     EMBL; AK052309; BAC34931.1; -.
DR
DR
     MGD; MGI:1860440; Bace2.
     GO; GO:0004194; F:pepsin A activity; IEA.
DR
     GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
     InterPro; IPR001969; Aspprotease_AS.
DR
     InterPro; IPR001461; Peptidase A1.
DR
     InterPro; IPR009007; Pept A acid.
DR
     Pfam; PF00026; asp; 1.
DR
     PRINTS; PRO0792; PEPSIN.
DR
     PROSITE; PS00141; ASP PROTEASE; 2.
DR
               514 AA; 55871 MW; 8BF45E07B0990225 CRC64;
SQ
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                         89.3%; Score 2399; DB 11; Length 514;
  Query Match
  Best Local Similarity 88.6%; Pred. No. 1e-178;
                                               35; Indels
                                                             4; Gaps
                                                                        1;
  Matches 459; Conservative 20; Mismatches
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Qу
              1 MGALLRALLLPVLAQWLLSAVPALAPAPFTLPLQVARATNHRASAVPGLGTPELPRADGL 60
Db
           61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qу
                          111111 1:
           61 ALALEPVRAT----ANFLAMVDNLQGDSGRGYYLEMLIGTPPQKVQILVDTGSSNFAVAG 116
Db
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121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qу
             117 APHSYIDTYFDSESSSTYHSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNSSFLVNIATI 176
Db
        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qy
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Db
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Qу
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Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qy
            297 IVDSGTTLLRLPQKVFDAVVEAVARTSLIPEFSDGFWTGAQLACWTNSETPWAYFPKISI 356
Db
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFD 420
Qy
            357 YLRDENASRSFRTTILPQLYIQPMMGAGFNYECYRFGISSSTNALVIGATVMEGFYVVFD 416
Db
        421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
Qу
            417 RAORRVGFAVSPCAEIEGTTVSEISGPFSTEDIASNCVPAQALNEPILWIVSYALMSVCG 476
Db
        481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qу
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ID
    09JL18
              PRELIMINARY;
                              PRT:
AC
    Q9JL18;
    01-OCT-2000 (TrEMBLrel. 15, Created)
DT
    01-OCT-2000 (TrEMBLrel. 15, Last sequence update)
DT
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DT
    Aspartyl protease 1.
DE
    BACE2.
GN
    Mus musculus (Mouse).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OC
    NCBI TaxID=10090;
OX
RN
    [1]
RP
    SEQUENCE FROM N.A.
    Choi D.K., Sugano S., Sakaki Y.;
RA
    "Molecular characterization of the mouse Aspl gene, a homolog of the
RT
    human ASP1 (Down Syndrome Region aspartyl protease).";
RT
    Submitted (DEC-1999) to the EMBL/GenBank/DDBJ databases.
RL
    -!- SIMILARITY: BELONGS TO PEPTIDASE FAMILY A1.
CC
    EMBL; AF216310; AAF36599.1; -.
DR
    HSSP; P00797; 2REN.
DR
    MEROPS; A01.041; -.
DR
    MGD; MGI:1860440; Bace2.
DR
    GO; GO:0004194; F:pepsin A activity; IEA.
DR
    GO; GO:0008233; F:peptidase activity; IEA.
DR
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
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InterPro; IPR001461; Peptidase A1.
DR
    InterPro; IPR009007; Pept A acid.
DR
    Pfam; PF00026; asp; 1.
DR
    PRINTS; PR00792; PEPSIN.
DR
    PROSITE; PS00141; ASP PROTEASE; 2.
DR
    Aspartyl protease; Hydrolase; Protease.
KW
             514 AA; 55799 MW; A70725F2C1DF5B47 CRC64;
    SEQUENCE
SO
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                                            Length 514;
 Query Match
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 Best Local Similarity
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Qу
                      61 ALALEPVRAT----ANFLAMVDNLQGDSGRGYYLEMLIGTPPQKVQILVDTGSSNFAVAG 116
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        121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qy
            117 APHSYIDTYFDSESSSTYHSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNSSFLVNIATI 176
Db
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Qу
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Db
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Qy
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Qy
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Db
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Qy
           357 YLRDENASRSFRITILPOLYIOPMMGAGFNYECYRFGISSSTNALVIGATVMEGFYVVFD 416
Db
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Db
        481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qу
           111111:1111 1: 111111111111111111
        477 AILLVLILLLLPLHCRHAPRDPEVVNDESSLVRHRWK 514
Db
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O9NZL2
                            PRT:
                                  468 AA.
    Q9NZL2
              PRELIMINARY;
ID
AC
    O9NZL2:
    01-OCT-2000 (TrEMBLrel. 15, Created)
DT
    01-OCT-2000 (TrEMBLrel. 15, Last sequence update)
DT
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DT
DE
    Aspartyl protease.
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GN
    BACE2.
    Homo sapiens (Human).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OC
OX
    NCBI TaxID=9606;
    [1]
RN
RP
    SEQUENCE FROM N.A.
    MEDLINE=20422477; PubMed=10965118;
RX
    Solans A., Estivill X., de La Luna S.;
RA
    "A new aspartyl protease on 21q22.3, BACE2, is highly similar to
RT
    Alzheimer's amyloid precursor protein beta-secretase.";
RT
    Cytogenet. Cell Genet. 89:177-184(2000).
RL
    -!- SIMILARITY: BELONGS TO PEPTIDASE FAMILY A1.
CC
    EMBL; AF188276; AAF35835.1; -.
DR
    HSSP; P00797; 2REN.
DR
    GO; GO:0004194; F:pepsin A activity; IEA.
DR
    GO; GO:0008233; F:peptidase activity; IEA.
DR
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR001461; Peptidase A1.
DR
    InterPro; IPR009007; Pept A acid.
DR
    Pfam; PF00026; asp; 1.
DR
DR
    PRINTS; PR00792; PEPSIN.
    PROSITE: PS00141; ASP PROTEASE; 2.
DR
    Aspartyl protease; Hydrolase; Protease.
KW
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SQ
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                      88.4%; Score 2375; DB 4; Length 468;
 Query Match
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 Best Local Similarity
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 Matches 468; Conservative
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Qу
            1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Db
         61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qy
            61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Dh
        121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qу
            121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
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        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qу
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Db
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qу
            241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Db
         301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qу
            301 IVDSGTTLLRLPQKVFDAVVEAVARASL----- 328
Db
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Qу
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Qу
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Db
         481 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 518
Qу
            431 AILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRHRWK 468
Db
RESULT 5
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               PRELIMINARY:
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ID
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AC
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    01-MAR-2001 (TrEMBLrel. 16, Created)
DT
    01-MAR-2001 (TrEMBLrel. 16, Last sequence update)
DT
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DΨ
DE
    CDA13.
OS
    Homo sapiens (Human).
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OC
    NCBI TaxID=9606;
ΟX
RN
    [1]
RP
    SEOUENCE FROM N.A.
RC
    TISSUE=Pheochromocytoma;
    Li Y., Huang Q., Peng, y, Song H., Yu Y., Xu S., Ren S., Chen Z.,
RA
RA
    Han Z.;
    Submitted (DEC-1999) to the EMBL/GenBank/DDBJ databases.
RL
    -!- SIMILARITY: BELONGS TO PEPTIDASE FAMILY A1.
CC
    EMBL; AF212252; AAG41783.1; -.
DR
    HSSP; P00797; 2REN.
DR
    GO; GO:0004194; F:pepsin A activity; IEA.
DR
ĎR
    GO; GO:0008233; F:peptidase activity; IEA.
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR001461; Peptidase A1.
DR
    InterPro; IPR009007; Pept A acid.
DR
    Pfam; PF00026; asp; 1.
DR
    PRINTS; PR00792; PEPSIN.
DR
    PROSITE; PS00141; ASP PROTEASE; 2.
DR
    Aspartyl protease; Hydrolase; Protease.
KW
    SEQUENCE 439 AA; 48275 MW; 02EC0E0E50F11602 CRC64;
SO
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Qу
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121 AYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVAGSGTNGGSLVLGGIEPSLY 180
Db
        260 KGDIWYTPIKEEWYYOIEILKLEIGGOSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAV 319
Qу
            181 KGDIWYTPIKEEWYYOIEILKLEIGGOSLNLDCREYNADKAIVDSGTTLLRLPOKVFDAV 240
Db
        320 VEAVARASLIPEFSDGFWTGSOLACWTNSETPWSYFPKISIYLRDENSSRSFRITILPOL 379
Qу
            241 VEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITILPQL 300
Db
        380 YIOPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFDRAOKRVGFAASPCAEIAGA 439
Qу
            Db
        301 YIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAASPCAEIAGA 360
        440 AVSEISGPFSTEDVASNCVPAOSLSEPILWIVSYALMSVCGAILLVLIVLLLLPFRCORR 499
Qу
            361 AVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCGAILLVLIVLLLLPFRCQRR 420
Db
        500 PRDPEVVNDESSLVRHRWK 518
Qу
            421 PRDPEVVNDESSLVRHRWK 439
Db
RESULT 6
Q8N2D4
ID
    Q8N2D4
              PRELIMINARY;
                              PRT;
                                    423 AA.
AC
    08N2D4;
    01-OCT-2002 (TrEMBLrel. 22, Created)
DT
    01-OCT-2002 (TrEMBLrel. 22, Last sequence update)
DТ
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DT
    Hypothetical protein OVARC1000363.
DE
OS
    Homo sapiens (Human).
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OC
OX
    NCBI TaxID=9606;
RN
    [1]
    SEQUENCE FROM N.A.
RΡ
    TISSUE=Ovarian carcinoma;
RC
RA
    Ota T., Nishikawa T., Suzuki Y., Kawai-Hio Y., Hayashi K., Ishii S.,
    Saito K., Yamamoto J., Wakamatsu A., Nagai T., Nakamura Y.,
RA
RA
    Nagahari K., Sugano S., Isogai T.;
    "HRI human cDNA sequencing project.";
RT
RL
    Submitted (MAR-2002) to the EMBL/GenBank/DDBJ databases.
DR
    EMBL; AK075539; BAC11682.1; -.
DR
    GO; GO:0004194; F:pepsin A activity; IEA.
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR001461; Peptidase Al.
    InterPro; IPR009007; Pept A acid.
DR
DR
    Pfam; PF00026; asp; 1.
DR
    PRINTS; PR00792; PEPSIN.
    PROSITE; PS00141; ASP PROTEASE; 2.
DR
KW
    Hypothetical protein.
SQ
    SEQUENCE
             423 AA; 46457 MW; 4D4839F2ED9C2CE1 CRC64;
                      81.3%; Score 2184; DB 4; Length 423;
 Query Match
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Best Local Similarity 99.3%; Pred. No. 4.7e-162;
                                                                0;
                          1; Mismatches
                                          2; Indels
                                                         Gaps
 Matches 420; Conservative
         96 MLIGTPPQKLQILVDTGSSNFAVAGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWT 155
Qу
           1 MLIGTPPQKLQILVDTGSSNFAVAGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWT 60
Dh
        156 GFVGEDLVTIPKGFNTSFLVNIATIFESENFFLPGIKWNGILGLAYATLAKPSSSLETFF 215
Qу
           61 GFVGEDLVTIPKGFNTSFLVNIATIFESGNFFLPGIQWNGILGLAYATLAKPSSSLETFF 120
Db
        216 DSLVTQANIPNVFSMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQ 275
Qy
           121 DSLVTOANIPNVFSMOMRGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQ 180
Db
        276 IEILKLEIGGOSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDG 335
Qy
           181 IEILKLEIGGOSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDG 240
Db
        336 FWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGLNYECYR 395
Qy
           241 FWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGLNYECYR 300
Db
        396 FGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVAS 455
Qy
           301 FGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVAS 360
Db
        456 NCVPAQSLSEPILWIVSYALMSVCGAILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRH 515
Qy
           361 NCVPAOSLSEPILWIVSYALMSVCGAILLVLIVLLLLPFRCQRRPRDPEVVNDESSLVRH 420
Db
        516 RWK 518
Qу
           IIII
Db
        421 RWK 423
RESULT 7
O9NZL1
              PRELIMINARY;
                             PRT;
                                   396 AA.
ID
    Q9NZL1
AC
    Q9NZL1;
    01-OCT-2000 (TrEMBLrel. 15, Created)
DT
    01-OCT-2000 (TrEMBLrel. 15, Last sequence update)
DΨ
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DT
    Aspartyl protease.
DΕ
    BACE2.
GN
    Homo sapiens (Human).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC.
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OC
    NCBI TaxID=9606;
OX
RN
    [1]
    SEOUENCE FROM N.A.
RP
    MEDLINE=20422477; PubMed=10965118;
RX
    Solans A., Estivill X., de La Luna S.;
RA
    "A new aspartyl protease on 21q22.3, BACE2, is highly similar to
RT
    Alzheimer's amyloid precursor protein beta-secretase.";
RТ
    Cytogenet. Cell Genet. 89:177-184(2000).
RL
    -!- SIMILARITY: BELONGS TO PEPTIDASE FAMILY A1.
CC
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DR
    EMBL; AF188277; AAF35836.1; -.
    HSSP; P00797; 2REN.
DR
    GO; GO:0004194; F:pepsin A activity; IEA.
DR
    GO; GO:0008233; F:peptidase activity; IEA.
DR
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR001461; Peptidase A1.
DR
DR
    InterPro; IPR009007; Pept A acid.
    Pfam; PF00026; asp; 1.
DR
DR
    PRINTS; PR00792; PEPSIN.
    PROSITE; PS00141; ASP PROTEASE; 2.
DR
KW
    Aspartyl protease; Hydrolase; Protease.
SO
    SEOUENCE
            396 AA;
                    43013 MW; 5023A7AF391CEAC9 CRC64;
                      73.2%; Score 1966; DB 4; Length 396;
 Query Match
 Best Local Similarity
                      100.0%; Pred. No. 4.5e-145;
 Matches 378; Conservative
                          0; Mismatches
                                           0; Indels
                                                       0;
                                                                 0;
                                                          Gaps
          1 MGALARALLLPLLAOWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qу
            1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Db
         61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qу
            61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Db
        121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Qy
            121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTOGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
Db
        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
Qv
            Db
        181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qy
            Db
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYOIEILKLEIGGOSLNLDCREYNADKA 300
Qy
        301 IVDSGTTLLRLPOKVFDAVVEAVARASLIPEFSDGFWTGSOLACWTNSETPWSYFPKISI 360
            Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qу
        361 YLRDENSSRSFRITILPQ 378
            Db
        361 YLRDENSSRSFRITILPQ 378
RESULT 8
07T0Y2
ID
    Q7T0Y2
              PRELIMINARY;
                             PRT;
                                   500 AA.
AC
    Q7T0Y2;
DT
    01-OCT-2003 (TrEMBLrel. 25, Created)
    01-OCT-2003 (TrEMBLrel. 25, Last sequence update)
DT
DT
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DΕ
    Hypothetical protein.
    Xenopus laevis (African clawed frog).
OS
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
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Amphibia; Batrachia; Anura; Mesobatrachia; Pipoidea; Pipidae;
OC
OC
    Xenopodinae; Xenopus.
OX
    NCBI TaxID=8355;
RN
    [1]
RP
    SEQUENCE FROM N.A.
RC
    TISSUE=Embryo;
    MEDLINE=22341132; PubMed=12454917;
RX
    Klein S.L., Strausberg R.L., Wagner L., Pontius J., Clifton S.W.,
RA
RA
    Richardson P.;
    "Genetic and genomic tools for Xenopus research: The NIH Xenopus
RT
RT
    initiative.";
RL
    Dev. Dyn. 225:384-391(2002).
RN
    [2]
    SEQUENCE FROM N.A.
RP
RC
    TISSUE=Embryo;
RX
    MEDLINE=22388257; PubMed=12477932;
    Strausberg R.L., Feingold E.A., Grouse L.H., Derge J.G.,
RA
    Klausner R.D., Collins F.S., Wagner L., Shenmen C.M., Schuler G.D.,
RA
    Altschul S.F., Zeeberg B., Buetow K.H., Schaefer C.F., Bhat N.K.,
RA
    Hopkins R.F., Jordan H., Moore T., Max S.I., Wang J., Hsieh F.,
RA
    Diatchenko L., Marusina K., Farmer A.A., Rubin G.M., Hong L.,
RA
    Stapleton M., Soares M.B., Bonaldo M.F., Casavant T.L., Scheetz T.E.,
RA
    Brownstein M.J., Usdin T.B., Toshiyuki S., Carninci P., Prange C.,
RA
    Raha S.S., Loquellano N.A., Peters G.J., Abramson R.D., Mullahy S.J.,
RA
    Bosak S.A., McEwan P.J., McKernan K.J., Malek J.A., Gunaratne P.H.,
RA
    Richards S., Worley K.C., Hale S., Garcia A.M., Gay L.J., Hulyk S.W.,
RA
    Villalon D.K., Muzny D.M., Sodergren E.J., Lu X., Gibbs R.A.,
RΑ
     Fahey J., Helton E., Ketteman M., Madan A., Rodrigues S., Sanchez A.,
RA
    Whiting M., Madan A., Young A.C., Shevchenko Y., Bouffard G.G.,
RA
    Blakesley R.W., Touchman J.W., Green E.D., Dickson M.C.,
RA
    Rodriguez A.C., Grimwood J., Schmutz J., Myers R.M., Butterfield Y.S.,
RA
     Krzywinski M.I., Skalska U., Smailus D.E., Schnerch A., Schein J.E.,
RA
RA
     Jones S.J., Marra M.A.;
     "Generation and initial analysis of more than 15,000 full-length human
RT
     and mouse cDNA sequences.";
RT
     Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903(2002).
RL
RN
     SEQUENCE FROM N.A.
RΡ
     TISSUE=Embryo;
RC
     Klein S., Strausberg R.;
RA
     Submitted (AUG-2003) to the EMBL/GenBank/DDBJ databases.
RL
     EMBL; BC055989; AAH55989.1; -.
DR
KW
     Hypothetical protein.
                       54722 MW; 10F16756CAFDCD0B CRC64;
     SEQUENCE
               500 AA;
SQ
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  Query Match
                         63.6%; Pred. No. 1.4e-123;
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  Matches 322; Conservative 74; Mismatches 100;
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                                                      Indels
                                                              10;
                                                                   Gaps
           13 LAOWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGLALALEPALASPA 72
Qу
                        11 11
            5 LVRLLLLCAAACASNKFIVPLNVSPAEIKGTLPV-APATPKDK--PGLLLA----SDPG 56
Db
           73 GAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGTPHSYIDTYFDT 132
Qу
             57 GTINFFSMVDNLAGDSGRGYYLELLIGSPPQKVNILVDTGSSNFAVAGSPNPDVNTFFDS 116
Db
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133 ERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFESENFFLPGIK 192
Qу
                     : |::|:|
        117 KLSTSYOSLNTEVTVRYTOGSWTGLLGKDVVSIPKGVNGTFLINIASIFOSESFFLPNIN 176
Db
        193 WNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVAGSGTNGGSLVLG 252
Qy
            1:1 1 11111
Db
        177 WQGILGLAYSTLAKPSSSVEPFFDSLVQQENIPDVFSMQMCGAGQSSPGNGINAGSLVLG 236
        253 GIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLP 312
QУ
            Db
        237 GVEPSLYKGNIWYTPITEEWYYQVEVLKFEVGGQRLNLDCTVYNSDKAIVDSGTTLLRLP 296
        313 QKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFR 372
Qv
             297 DKVFNAMVDAIVQTSLIQNFNAEFWAGLQLACWDKTQQPWNYFPDISIYLRDTNTSRSFR 356
Db
        373 ITILPQLYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAASP 432
Qy
            357 LTLKPQLYIQSVLTFQESLNCFRFGISQSASTLVIGATVMEGFYVIFDRAEKRVGFAVSS 416
Db
        433 CAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCGAILLVLIVLLLL 492
Qy
            |||::| ||||:||| | ||:|||:
                                     417 CAEVSGITVSEIAGPFGTSDVSSNCIARNPLREPIMWIISYSLMSLCGMILLVLVILLLL 476
Db
        493 PFRCQRRPRDPEVVNDESSLVRHRWK 518
Qv
              | :| | :||||||:||
        477 SNR--ORHDDMETINDESSLVOHRWK 500
Db
RESULT 9
09R1P7
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                               PRT:
                                     255 AA.
ID
    Q9R1P7
AC
    Q9R1P7;
    01-MAY-2000 (TrEMBLrel. 13, Created)
DT
    01-MAY-2000 (TrEMBLrel. 13, Last sequence update)
DT
DТ
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DE
    Aspartyl protease (Fragment).
GN
    BACE2.
OS
    Mus musculus (Mouse).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OX
    NCBI TaxID=10090;
RN
    [1]
RΡ
    SEQUENCE FROM N.A.
    Accarino M., Fumagalli P., Taramelli R., Ottolenghi S.;
RA
    "Cloning of a gene from chromosome 21 Down Region encoding a potential
RT
    transmembrane protease.";
RT
RL
    Submitted (FEB-1998) to the EMBL/GenBank/DDBJ databases.
    EMBL; AF051150; AAD45964.1; -.
DR
    MEROPS; A01.041; -.
DR
    MGD; MGI:1860440; Bace2.
DR
DR
    GO; GO:0004190; F:aspartic-type endopeptidase activity; IEA.
    GO; GO:0008233; F:peptidase activity; IEA.
DR
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR009007; Pept A acid.
DR
    PROSITE; PS00141; ASP PROTEASE; 1.
DR
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KW
    Protease.
FT
    NON TER
SQ
    SEQUENCE
              255 AA;
                       28685 MW; 53DE317815996D63 CRC64;
 Query Match
                        46.4%; Score 1246; DB 11;
                                                  Length 255;
 Best Local Similarity
                        91.0%; Pred. No. 4e-89;
 Matches 232; Conservative
                            12: Mismatches
                                              11;
                                                            0;
                                                               Gaps
                                                                       0;
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         264 WYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAV 323
Qy
             Db
           1 WYTPIKEEWYYQIEILKLEIGGQNLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAV 60
         324 ARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITILPQLYIQP 383
Qy
             61 ARTSLIPEFSDGFWTGAQLACWTNSETPWAYFPKISIYLRDENASRSFRITILPQLYIQP 120
Db
         384 MMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAASPCAEIAGAAVSE 443
Qy
             121 MMGAGFNYECYRFGISSSTNALVIGATVMEGFYVVFDRAQRRVGFAVSPCAEIEGTTVSE 180
Db
         444 ISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCGAILLVLIVLLLLPFRCQRRPRDP 503
Qy
             181 ISGPFSTEDIASNCVPAQALNEPILWIVSYALMSVCGAILLVLILLLLVPLHCRHAPRDP 240
Db
         504 EVVNDESSLVRHRWK 518
Qγ
             111111111111111
         241 EVVNDESSLVRHRWK 255
Db
RESULT 10
08C7R1
TD
               PRELIMINARY;
                                PRT:
                                       501 AA.
    Q8C7R1
AC
    Q8C7R1;
    01-MAR-2003 (TrEMBLrel. 23, Created)
01-MAR-2003 (TrEMBLrel. 23, Last sequence update)
DT
DT
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DT
    Beta-site APP cleaving enzyme.
DE
GN
    BACE.
OS
    Mus musculus (Mouse).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC.
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OX
    NCBI_TaxID=10090;
RN
    [1]
RP
    SEQUENCE FROM N.A.
RC
    STRAIN=C57BL/6J; TISSUE=Spinal cord;
RX
    MEDLINE=22354683; PubMed=12466851;
    The FANTOM Consortium,
RA
RA
    the RIKEN Genome Exploration Research Group Phase I & II Team;
RT
    "Analysis of the mouse transcriptome based on functional annotation of
    60,770 full-length cDNAs.";
RT
    Nature 420:563-573(2002).
RT.
DR
    EMBL; AK049626; BAC33844.1; -.
DR
    MGD; MGI:1346542; Bace.
    GO; GO:0004194; F:pepsin A activity; IEA.
DR
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR001461; Peptidase A1.
DR
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DR
     InterPro; IPR009007; Pept A acid.
DR
     Pfam; PF00026; asp; 1.
     PRINTS; PR00792; PEPSIN.
DR
DR
     PROSITE; PS00141; ASP PROTEASE; 1.
SO
              501 AA; 55761 MW; B410DA8B64647663 CRC64;
                      44.1%; Score 1186; DB 11; Length 501;
  Query Match
  Best Local Similarity 46.2%; Pred. No. 5.4e-84;
  Matches 238; Conservative 82; Mismatches 169; Indels
                                                       26; Gaps
                                                                 7:
           9 LLPLLAQWLLRAAPELAPAPFT-----LPLRVAAATNRVVAPTPGPGTPAERHADGLALA 63
Qу
                                  : | | | | | | | |
           1 MAPALHWLLLWVGSGMLPAOGTHLGIRLPLRSGLA----GPPLGLRLPRETDEES---- 51
Db
          64 LEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGTPH 123
Qy
                  52 ----EEPGRRGSFVEMVDNLRGKSGQGYYVEMTIGSPPQTLNILVDTGSSNFAVGAAPH 106
Db
         124 SYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFES 183
Qy
                              :: 1: : 11111
         107 PFLHRYYQRQLSSTYRDLRKGVYVPYTQGKWEGELGTDLVSIPHGPNVTVRANIAAITES 166
. Db
         184 ENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPV---A 240
Qy
            167 DKFFINGSNWEGILGLAYAEIARPDDSLEPFFDSLVKOTHIPNIFSLQLCGAGFPLNQTE 226
Db
         241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qу
               227 ALASVGGSMIIGGIDHSLYTGSLWYTPIRREWYYEVIIVRVEINGQDLKMDCKEYNYDKS 286
Db
         301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qу
            287 IVDSGTTNLRLPKKVFEAAVKSIKAASSTEKFPDGFWLGEQLVCWQAGTTPWNIFPVISL 346
Db
         361 YLRDENSSRSFRITILPQLYIQPMMGAGLNY-ECYRFGISPSTNALVIGATVMEGFYVIF 419
Qу
            347 YLMGEVTNQSFRITILPQQYLRPVEDVATSQDDCYKFAVSQSSTGTVMGAVIMEGFYVVF 406
Db
         420 DRAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVC 479
Qy
                               :: | | | | | | | | | |
                                                   | | ::| : ::|
             |||:||:||| |
         407 DRARKRIGFAVSACHVHDEFRTAAVEGPFVTADMEDCGYNIPQTDESTLMTIAYVMAAIC 466
Db
         480 GAILLVLIVLLLLPFRCQR--RPRDPEVVNDESSL 512
Qу
             ]: :: : |:: :|| | | : : : | | | |
         467 -ALFMLPLCLMVCQWRCLRCLRHQHDDFADDISLL 500
Db
RESULT 11
O9ULS1
                              PRT;
                                    532 AA.
ID
     O9ULS1
               PRELIMINARY;
     09ULS1;
AC
     01-MAY-2000 (TrEMBLrel. 13, Created)
DT
     01-OCT-2001 (TrEMBLrel. 18, Last sequence update)
DT
     01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
דת
     Hypothetical protein KIAA1149 (Fragment).
DF.
 GN
     KIAA1149.
 OS
     Homo sapiens (Human).
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Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
        Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OC
        NCBI TaxID=9606;
OX
RN
RP
        SEQUENCE FROM N.A.
RC
        TISSUE=Brain:
        MEDLINE=20039618; PubMed=10574461;
RX
        Hirosawa M., Nagase T., Ishikawa K., Kikuno R., Nomura N., Ohara O.;
RA
        "Characterization of cDNA clones selected by the GeneMark analysis
RT
RТ
        from size-fractionated cDNA libraries from human brain.";
        DNA Res. 6:329-336(1999).
RL
        -!- SIMILARITY: BELONGS TO PEPTIDASE FAMILY A1.
CC
        EMBL; AB032975; BAA86463.2; -.
DR
        HSSP; P56272; 1AM5.
DR
        GO; GO:0004194; F:pepsin A activity; IEA.
DR
        GO; GO:0008233; F:peptidase activity; IEA.
DR
        GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
        InterPro; IPR001969; Aspprotease AS.
DR
        InterPro; IPR001461; Peptidase Al.
DR
        InterPro; IPR009007; Pept A acid.
DR
        Pfam; PF00026; asp; 1.
DR
DR
        PRINTS; PR00792; PEPSIN.
        PROSITE; PS00141; ASP PROTEASE; 1.
DR
        Hypothetical protein; Aspartyl protease; Hydrolase; Protease.
KW
FT
        NON TER
                               1
        SEQUENCE
                           532 AA: 58720 MW; 98B135D0D5FBD2E8 CRC64;
SO
                                            44.0%; Score 1183.5; DB 4; Length 532;
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   Matches 231; Conservative 81; Mismatches 154; Indels
                                                                                                              17: Gaps
                                                                                                                                    6;
                   44 APTPGPGTPAERHADGLALAL-----EPALASPAGAANFLAMVDNLQGDSGRGYYLE 95
Qу
                                          1 1 1
                                                                                           :|: |||||:| ||:||
                        11: 11
                                                                         52 APSTASGCPCAAAWGGAPLGLRLPRETDEEP--EEPGRRGSFVEMVDNLRGKSGQGYYVE 109
Db
                   96 MLIGTPPOKLOILVDTGSSNFAVAGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWT 155
Qy
                        1:1:111 | 1111111111 | 11:: 1: : | 11| | 11
                                                                                                           Db
                 110 MTVGSPPQTLNILVDTGSSNFAVGAAPHPFLHRYYQRQLSSTYRDLRKGVYVPYTQGKWE 169
                 156 GFVGEDLVTIPKGFNTSFLVNIATIFESENFFLPGIKWNGILGLAYATLAKPSSSLETFF 215
Qу
                        170 GELGTDLVSIPHGPNVTVRANIAAITESDKFFINGSNWEGILGLAYAEIARPDDSLEPFF 229
Db
                 216 DSLVTQANIPNVFSMQMCGAGLPVAGS---GTNGGSLVLGGIEPSLYKGDIWYTPIKEEW 272
Qу
                        1111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 111 | 
                 230 DSLVKQTHVPNLFSLQLCGAGFPLNQSEVLASVGGSMIIGGIDHSLYTGSLWYTPIRREW 289
Db
                 273 YYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEF 332
Qу
                        290 YYEVIIVRVEINGQDLKMDCKEYNYDKSIVDSGTTNLRLPKKVFEAAVKSIKAASSTEKF 349
Db
                 333 SDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGLNY- 391
Qу
                                                    350 PDGFWLGEOLVCWOAGTTPWNIFPVISLYLMGEVTNOSFRITILPQQYLRPVEDVATSQD 409
Db
                 392 ECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAASPCAEIAGAAVSEISGPFSTE 451
Qy
                        :||:| || ||: ||:|| :|||||:|||:||:||| ||:||
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410 DCYKFAISQSSTGTVMGAVIMEGFYVVFDRARKRIGFAVSACHVHDEFRTAAVEGPFVTL 469
Db
         452 DVASNCVPAQSLSEPILWIVSYALMSVCGAILLVLIVLLLLPFRCQR--RPRDPEVVNDE 509
Qy
                          | | ::|:::|::::|::::|:::::|
         470 DMEDCGYNIPQTDESTLMTIAYVMAAIC-ALFMLPLCLMVCQWRCLRCLRQQHDDFADDI 528
Db
         510 SSL 512
Qy
             Db
         529 SLL 531
RESULT 12
Q8BQY4
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                                  PRT;
                                        501 AA.
ΙD
    O8BOY4
AC
    O8BOY4;
    01-MAR-2003 (TrEMBLrel. 23, Created)
DΤ
    01-MAR-2003 (TrEMBLrel. 23, Last sequence update)
DT
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DT
DE
    Beta-site APP cleaving enzyme.
GN
    BACE.
OS
    Mus musculus (Mouse).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OC
OX
    NCBI TaxID=10090;
RN
    [1]
    SEQUENCE FROM N.A.
RP
    STRAIN=C57BL/6J; TISSUE=Brain;
RC
    MEDLINE=22354683; PubMed=12466851;
RX
    The FANTOM Consortium,
RA
    the RIKEN Genome Exploration Research Group Phase I & II Team;
RA
    "Analysis of the mouse transcriptome based on functional annotation of
RT
RT
     60,770 full-length cDNAs.";
    Nature 420:563-573(2002).
RL
    EMBL; AK046175; BAC32620.1; -.
DR
DR
    MGD; MGI:1346542; Bace.
    GO; GO:0004194; F:pepsin A activity; IEA.
DR
DR
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
    InterPro; IPR001969; Aspprotease AS.
DR
DR
    InterPro; IPR001461; Peptidase A1.
DR
    InterPro; IPR009007; Pept A acid.
DR
    Pfam; PF00026; asp; 1.
DR
    PRINTS; PR00792; PEPSIN.
DR
    PROSITE; PS00141; ASP PROTEASE; 1.
SO
               501 AA; 55816 MW; C0855513145E024E CRC64;
 Query Match
                         44.0%; Score 1183; DB 11;
                                                     Length 501;
 Best Local Similarity
                         46.0%; Pred. No. 9.2e-84;
 Matches 237; Conservative 83; Mismatches 169; Indels
                                                              26; Gaps
                                                                          7;
           9 LLPLLAQWLLRAAPELAPAPFT----LPLRVAAATNRVVAPTPGPGTPAERHADGLALA 63
Qy
                           : || |
                                    - 11
Db
           1 MAPALHWLLLWVGSGMLPAQGTHLGIRLPLRSGLA-----GPPLGLRLPRETDEES---- 51
          64 LEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGTPH 123
Qy
                        Db
          52 ----EEPGRRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFAVGAAPH 106
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124 SYIDTYFDTERSSTYRSKGFDVTVKYTOGSWTGFVGEDLVTIPKGFNTSFLVNIATIFES 183
Qy
                              :: |: : ||||
        107 PFLHRYYQRQLSSTYRDLRKGVYVPYTQGKWEGELGTDLVSIPHGPNVTVRANIAAITES 166
Db
        184 ENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTOANIPNVFSMOMCGAGLPV---A 240
Qу
            167 DKFFINGSNWEGILGLAYAEIARPDDSLEPFFDSLVKOTHIPNIFSLOLCGAGFPLNQTE 226
Db
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qy
               227 ALASVGGSMIIGGIDHSLYTGRLWYTPIRREWYYEVIIVRVEINGODLKMDCKEYNYDKS 286
Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
QУ
            287 IVDSGTTNLRLPKKVFEAAVKSIKAASSTEKFPDGFWLGEOLVCWOAGTTPWNIFPVISL 346
Db
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNY-ECYRFGISPSTNALVIGATVMEGFYVIF 419
Qу
            347 YLMGEVTNQSFRITILPQQYLRPVEDVATSQDDCYKFAVSQSSTGTVMGAVIMEGFYVVF 406
Db
        420 DRAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVC 479
Qу
            |||:||:||| |
                               :: ||| | |:
                                                    | | ::| : ::|
        407 DRARKRIGFAVSACHVHDEFRTAAVEGPFVTADMEDCGYNIPQTDESTLMTIAYVMAAIC 466
Db
        480 GAILLVLIVLLLLPFRCQR--RPRDPEVVNDESSL 512
Qу
             1: :: : 1:: : : : : : : : : | | |
        467 -ALFMLPLCLMVCQWRCLRCLRHQHDDFADDISLL 500
Db
RESULT 13
08IYC8
ΙD
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               PRELIMINARY:
                               PRT:
                                     501 AA.
AC
    O8IYC8;
    01-MAR-2003 (TrEMBLrel. 23, Created)
DT
    01-MAR-2003 (TrEMBLrel. 23, Last sequence update)
DT
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DT
DE
    Beta-site APP-cleaving enzyme.
OS
    Homo sapiens (Human).
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
OC
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OX
    NCBI TaxID=9606;
RN
    [1]
RP
    SEQUENCE FROM N.A.
RC
    TISSUE=Brain;
RA
    Strausberg R.;
    Submitted (JUL-2002) to the EMBL/GenBank/DDBJ databases.
RL
DR
    EMBL; BC036084; AAH36084.1; -.
DR
    GO; GO:0004194; F:pepsin A activity; IEA.
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR001461; Peptidase A1.
DR
DR
    InterPro; IPR009007; Pept A acid.
DR
    Pfam; PF00026; asp; 1.
    PRINTS; PR00792; PEPSIN.
DR
    PROSITE; PS00141; ASP PROTEASE; 1.
DR
SO
    SEOUENCE
             501 AA; 55824 MW; 768595CF5517EFB7 CRC64;
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 Query Match
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 Matches 239; Conservative 82; Mismatches 165; Indels
                                                        33; Gaps 9;
          7 ALLLPLLAOWLLRAAPELAPAPFT----LPLRVAAATNRVVAPTPGPGTPAERHADGLA 61
Qy
                                     1111
                          : || |
          2 AOALPWLLLWM---GAGVLPAHGTOHGIRLPLRSGLG-----GAPL----GLR 42
Db
         62 LALE--PALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVA 119
Qу
            43 LPRETDEEPEEPGRRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFAVG 102
Db
        120 GTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIAT 179
Qy
              103 AAPHPFLHRYYOROLFSTYRDLRKGVYVPYTOGKWEGELGTDLVSIPHGPNVTVRANIAA 162
Db
        180 IFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPV 239
Qу
            163 ITESDKFFINGSNWEGILGLAYAEIARPDDSLEPFFDSLVKOTHVPNLFSLOLCGAGFPL 222
Db
        240 AGS---GTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYN 296
Qу
                 Db
        223 NQSEVLASVGGSMIIGGIDHSLYTGSLWYTPIRREWYYEVIIVRVEINGQDLKMDCKEYN 282
        297 ADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFP 356
Qу
             283 YDKSIVDSGTTNLRLPKKVFEAAVKSIKAASSTEKFPDGFWLGEQLVCWQAGTTPWNIFP 342
Db
        357 KISIYLRDENSSRSFRITILPOLYIOPMMGAGLNY-ECYRFGISPSTNALVIGATVMEGF 415
Qy
             11:11 1 :::111111111 1::1: : :11:1 11 1::
        343 VISLYLMGEVTNOSFRITILPOOYLRPVEDVATSODDCYKFAISOSSTGTVMGAVIMEGF 402
Db
        416 YVIFDRAOKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAOSLSEPILWIVSYAL 475
Qу
            11:||||:||:||| : : : ||| | | : :
                                                       | | ::|:
        403 YVVFDRARKRIGFAVSACHVHDEFRTAAVEGPFVTLDMEDCGYNIPOTDESTLMTIAYVM 462
Db
        476 MSVCGAILLVLIVLLLLPFRCOR--RPRDPEVVNDESSL 512
Qу
             ::| |: :: : |:: :|| | | : : :| | |
        463 AAIC-ALFMLPLCLMVCQWRCLRCLRQQHDDFADDISLL 500
Db
RESULT 14
Q8C4F4
ID
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                              PRT;
                                    467 AA.
AC
    08C4F4;
DT
    01-MAR-2003 (TrEMBLrel. 23, Created)
    01-MAR-2003 (TrEMBLrel. 23, Last sequence update)
ÐΤ
    01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DT
DE
    Beta-site APP cleaving enzyme.
GN
    BACE.
OS
    Mus musculus (Mouse).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OC
OX
    NCBI TaxID=10090;
RN
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RP
    SEQUENCE FROM N.A.
RC.
    STRAIN=C57BL/6J; TISSUE=Cerebellum;
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MEDLINE=22354683; PubMed=12466851;
RX
    The FANTOM Consortium,
RA
    the RIKEN Genome Exploration Research Group Phase I & II Team;
RA
    "Analysis of the mouse transcriptome based on functional annotation of
RT
    60,770 full-length cDNAs.";
RT
    Nature 420:563-573(2002).
RL
    EMBL; AK082317; BAC38462.1; -.
DR
    MGD; MGI:1346542; Bace.
DR
    GO; GO:0004194; F:pepsin A activity; IEA.
DR
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR001461; Peptidase Al.
DR
    InterPro; IPR009007; Pept A acid.
DR
    Pfam; PF00026; asp; 1.
DR
    PRINTS; PR00792; PEPSIN.
DR
    PROSITE; PS00141; ASP PROTEASE; 1.
DR
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SQ
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 Matches 215; Conservative 77; Mismatches 163; Indels
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                                                       60;
          9 LLPLLAQWLLRAAPELAPAPFT----LPLRVAAATNRVVAPTPGPGTPAERHADGLALA 63
Qy
            1 MAPALHWLLLWVGSGMLPAQGTHLGIRLPLRSGLA----GPPLGLRLPRETDEES---- 51
Db
         64 LEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGTPH 123
Qу
                      :1: |||||:|| ||:|||:|| :|:||| | ||:|||||
         52 ----EEPGRRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFAVGAAPH 106
Db
        124 SYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFES 183
Qу
             107 PFLHRYYOROLSSTYRDLRKGVYVPYTQGKWEGELGTDLVSIPHGPNVTVRANIAAITES 166
Db
        184 ENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPV---A 240
Qу
            167 DKFFINGSNWEGILGLAYAEIARPDDSLEPFFDSLVKQTHIPNIFSLQLCGAGFPLNQTE 226
Db,
        241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qу
               227 ALASVGGSMIIGGIDHSLYTGSLWYTPIRREWYYEVIIVRVEINGQDLKMDCKE----- 280
Db
        301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qу
                                      111: 11 11:
        281 -----TEKFPDGFWLGEQLVCWQAGTTPWNIFPVISL 312
Db
        361 YLRDENSSRSFRITILPQLYIQPMMGAGLNY-ECYRFGISPSTNALVIGATVMEGFYVIF 419
QУ
            313 YLMGEVTNQSFRITILPQQYLRPVEDVATSQDDCYKFAVSQSSTGTVMGAVIMEGFYVVF 372
Db
        420 DRAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVC 479
Qу
                              :: | | | | | | | | | |
                                                   1 | ::| : ::|
            373 DRARKRIGFAVSACHVHDEFRTAAVEGPFVTADMEDCGYNIPQTDESTLMTIAYVMAAIC 432
Db
         480 GAILLVLIVLLLLPFRCQR--RPRDPEVVNDESSL 512
Qy
             1: :: : |:: :|| | | : : :| | |
         433 -ALFMLPLCLMVCQWRCLRCLRHQHDDFADDISLL 466
Db
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RESULT 15
Q9CUU5
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ID
                  PRELIMINARY;
                                    PRT;
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AC
DT
     01-JUN-2001 (TrEMBLrel. 17, Created)
     01-MAR-2003 (TrEMBLrel. 23, Last sequence update)
DT
     01-OCT-2003 (TrEMBLrel. 25, Last annotation update)
DT
     Adult male brain cDNA, RIKEN full-length enriched library,
DE
DE
     clone: 3526402A15 product: beta-site APP cleaving enzyme, full insert
DE
     sequence (Fragment).
GN
     BACE.
OS
     Mus musculus (Mouse).
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
OC
     Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OX
     NCBI TaxID=10090;
RN
     [1]
     SEQUENCE FROM N.A.
RP
RC
     STRAIN=C57BL/6J; TISSUE=Brain;
RA
     Adachi J., Aizawa K., Akahira S., Akimura T., Arai A., Aono H.,
     Arakawa T., Bono H., Carninci P., Fukuda S., Fukunishi Y., Furuno M.,
RA
RA
     Hanagaki T., Hara A., Hayatsu N., Hiramoto K., Hiraoka T., Hori F.,
RA
     Imotani K., Ishii Y., Itoh M., Izawa M., Kasukawa T., Kato H.,
     Kawai J., Kojima Y., Konno H., Kouda M., Koya S., Kurihara C.,
RA
     Matsuyama T., Miyazaki A., Nishi K., Nomura K., Numazaki R., Ohno M.,
RA
RA
     Okazaki Y., Okido T., Owa C., Saito H., Saito R., Sakai C., Sakai K.,
     Sano H., Sasaki D., Shibata K., Shibata Y., Shinagawa A., Shiraki T.,
RA
RA
     Sogabe Y., Suzuki H., Tagami M., Tagawa A., Takahashi F., Tanaka T.,
RA
     Tejima Y., Toya T., Yamamura T., Yasunishi A., Yoshida K., Yoshino M.,
RA
     Muramatsu M., Hayashizaki Y.;
     Submitted (JUL-2000) to the EMBL/GenBank/DDBJ databases.
RL
RN
     [2]
RP
     SEQUENCE FROM N.A.
     STRAIN=C57BL/6J; TISSUE=Brain;
RC
RX
     MEDLINE=22354683; PubMed=12466851;
RA
     The FANTOM Consortium,
RA
     the RIKEN Genome Exploration Research Group Phase I & II Team;
RT
     "Analysis of the mouse transcriptome based on functional annotation of
RT
     60,770 full-length cDNAs.";
RL
     Nature 420:563-573(2002).
RN
     [3]
RP
     SEQUENCE FROM N.A.
RC
     STRAIN=C57BL/6J; TISSUE=Brain;
RX
     MEDLINE=21085660; PubMed=11217851;
RA
     RIKEN FANTOM Consortium;
     "Functional annotation of a full-length mouse cDNA collection.";
RT
RL
     Nature 409:685-690(2001).
RN
     [4]
     SEQUENCE FROM N.A.
RP
     STRAIN=C57BL/6J; TISSUE=Brain;
RC
RX
     MEDLINE=99279253; PubMed=10349636;
RA
     Carninci P., Hayashizaki Y.;
RT
     "High-efficiency full-length cDNA cloning.";
RL
     Meth. Enzymol. 303:19-44(1999).
RN
RP
     SEQUENCE FROM N.A.
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RC
     STRAIN=C57BL/6J; TISSUE=Brain;
RX
    MEDLINE=20499374; PubMed=11042159;
RA
     Carninci P., Shibata Y., Hayatsu N., Sugahara Y., Shibata K., Itoh M.,
RA
     Konno H., Okazaki Y., Muramatsu M., Hayashizaki Y.;
RT
     "Normalization and subtraction of cap-trapper-selected cDNAs to
RT
    prepare full-length cDNA libraries for rapid discovery of new genes.";
RL
    Genome Res. 10:1617-1630(2000).
RN
    SEQUENCE FROM N.A.
RP
RC
    STRAIN=C57BL/6J; TISSUE=Brain;
RX
    MEDLINE=20530913; PubMed=11076861;
RA
    Shibata K., Itoh M., Aizawa K., Nagaoka S., Sasaki N., Carninci P.,
    Konno H., Akiyama J., Nishi K., Kitsunai T., Tashiro H., Itoh M.,
RA
RA
    Sumi N., Ishii Y., Nakamura S., Hazama M., Nishine T., Harada A.,
RA
    Yamamoto R., Matsumoto H., Sakaguchi S., Ikegami T., Kashiwagi K.,
RA
    Fujiwake S., Inoue K., Togawa Y., Izawa M., Ohara E., Watahiki M.,
    Yoneda Y., Ishikawa T., Ozawa K., Tanaka T., Matsuura S., Kawai J.,
RA
    Okazaki Y., Muramatsu M., Inoue Y., Kira A., Hayashizaki Y.;
RA
RT
    "RIKEN integrated sequence analysis (RISA) system-384-format
RT
    sequencing pipeline with 384 multicapillary sequencer.";
RL
    Genome Res. 10:1757-1771(2000).
DR
    EMBL; AK014390; BAB29317.2; -.
DR
    MGD; MGI:1346542; Bace.
DR
    GO; GO:0004194; F:pepsin A activity; IEA.
DR
    GO; GO:0006508; P:proteolysis and peptidolysis; IEA.
DR
    InterPro; IPR001461; Peptidase A1.
DR
    InterPro; IPR009007; Pept A acid.
DR
    Pfam; PF00026; asp; 1.
FT
    NON TER
    SEQUENCE
SO
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 Query Match
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  Best Local Similarity
                        45.3%; Pred. No. 8.8e-43;
 Matches 121; Conservative 56; Mismatches
                                              86; Indels
                                                             4; Gaps
                                                                         3;
Qy
         249 LVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTL 308
             Db
           1 MIIGGIDHSLYTGSLWYTPIRREWYYEVIIVRVEINGQDLKMDCKEYNYDKSIVDSGTTN 60
Qy
         309 LRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSS 368
             :| |||| | || ||
                                                    Db
          61 LRLPKKVFEAAVKSIKAASSTEKFPDGFWLGEQLVCWQAGTTPWNIFPVISLYLMGEVTN 120
         369 RSFRITILPQLYIQPMMGAGLNY-ECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVG 427
Qy
             121 QSFRITILPQQYLRPVEDVATSQDDCYKFAVSQSSTGTVMGAVIMEGFYVVFDRARKRIG 180
Db
         428 FAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCGAILLVLI 487
Qy
                          :: | | | | | | :
                                                | | ::| : ::| |: :: :
Db
         181 FAVSACHVHDEFRTAAVEGPFVTADMEDCGYNIPQTDESTLMTIAYVMAAIC-ALFMLPL 239
         488 VLLLLPFRCQR--RPRDPEVVNDESSL 512
Qy
              1:: :|| | | : : :| | |
Db
         240 CLMVCQWRCLRCLRHQHDDFADDISLL 266
Search completed: March 4, 2004, 15:38:52
Job time : 76.3936 secs
```

## GenCore version 5.1.6 Copyright (c) 1993 - 2004 Compugen Ltd.

OM protein - protein search, using sw model

Run on: March 4, 2004, 15:22:30; Search time 16.5319 Seconds

(without alignments)

1631.532 Million cell updates/sec

ex legistr

Title: US-09-668-314C-2

Perfect score: 2687

Sequence: 1 MGALARALLLPLLAQWLLRA.....RPRDPEVVNDESSLVRHRWK 518

Scoring table: BLOSUM62

Gapop 10.0 , Gapext 0.5

Searched: 141681 seqs, 52070155 residues

Total number of hits satisfying chosen parameters: 141681

Minimum DB seq length: 0

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%

Listing first 45 summaries

Database: SwissProt 42:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

## **ŞUMMARIES**

		ક				
Result		Query				
No.	Score	Match	Length	DB	ID	Description
1	2687	100.0	518	1	BAE2 HUMAN	Q9y5z0 homo sapien
2	1187	44.2	501	1	BACE RAT	P56819 rattus norv
3	1185	44.1	501	1	BACE MOUSE	P56818 mus musculu
4	1178.5	43.9	501	1	BACE HUMAN	P56817 homo sapien
5	363.5	13.5	377	1	PEPC MACFU	P03955 macaca fusc
6	353	13.1	388	1	PEPC HUMAN	P20142 homo sapien
7	351.5	13.1	388	1	PEPC CALJA	Q9n2d3 callithrix
8	324.5	12.1	394	1	PEPC CAVPO	Q64411 cavia porce
9	320	11.9	402	1	RENI MOUSE	P06281 mus musculu
10	315.5	11.7	396	1	CATD CLUHA	Q9dex3 clupea hare
11	313.5	11.7	509	1	APR1 ORYSA	Q42456 oryza sativ
12	313	11.6	392	J 1	PEPC RAT	P04073 rattus norv
13	310	11.5	383	1	PEPE CHICK	P16476 gallus gall
14	308.5	11.5	412	1	CATD HUMAN	P07339 homo sapien
15	306.5	11.4	410	1	CATD MOUSE	P18242 mus musculu
16	305.5	11.4	401	1	RENS MOUSE	P00796 mus musculu
17	305	11.4	407	1	CATD RAT	P24268 rattus norv

18	302	11.2	324	1	PEP1 GADMO	P56272	gadus morhu
19	302	11.2	405	1	CARP YEAST	P07267	saccharomyc
20	301.5	11.2	398	1	CATE RAT	P16228	rattus norv
21	300.5	11.2	387	1	PEP2 RABIT	P27821	oryctolagus
22	300.5	11.2	397	1	CATE MOUSE	P70269	mus musculu
23	299	11.1	398	1	CATD CHICK	Q05744	gallus gall
24	298.5	11.1	387	1	PEP4 RABIT	P28713	oryctolagus
25	298.5	11.1	400	1	RENI_SHEEP	P52115	ovis aries
26	297	11.1	388	1	PEPA HUMAN	P00790	homo sapien
27	294.5	11.0	388	1	PEP2 MACFU	P27677	macaca fusc
28	291	10.8	388	1	PEP4 MACFU	P27678	macaca fusc
29	291	10.8	402	1	RENI_RAT	P08424	rattus norv
30	291	10.8	406	1	RENI_HUMAN	P00797	homo sapien
31	291	10.8	406	1	RENI_PANTR	P60016	pan troglod
32	290.5	10.8	396	1	CATE RABIT	P43159	oryctolagus
33	289	10.8	387	1	PEP3_RABIT	P27822	oryctolagus
34	289	10.8	388	1	PAG_HORSE	Q28389	equus cabal
35	288.5	10.7	390	1	CATD_BOVIN	P80209	bos taurus
36	288	10.7	387	1	PEP1_RABIT	P28712	oryctolagus
37	288	10.7	388	1	PEP1_MACFU	P03954	macaca fusc
38	287	10.7	367	1	PEPA_CHICK	P00793	gallus gall
39	287	10.7	391	1	CATE_CAVPO	P25796	cavia porce
40	287	10.7	396	1	CATE_HUMAN	P14091	homo sapien
41	286	10.6	388	1	PEPA_MACMU	P11489	macaca mula
42	285.5	10.6	387	1	PEPA_CALJA	Q9n2d4	callithrix
43	285	10.6	396	1	CARP_NEUCR	Q01294	neurospora
44	284.5	10.6	386	1	PEPA_PIG		sus scrofa
45	283	10.5	388	1	PEPF_RABIT	P27823	oryctolagus

## ALIGNMENTS

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RESULT 1
BAE2 HUMAN
ID
     BAE2 HUMAN
                    STANDARD;
                                   PRT;
                                           518 AA.
AC
     Q9Y5Z0; Q9UJT6;
DT
     30-MAY-2000 (Rel. 39, Created)
DT
     30-MAY-2000 (Rel. 39, Last sequence update)
DT
     10-OCT-2003 (Rel. 42, Last annotation update)
DE
     Beta secretase 2 precursor (EC 3.4.23.45) (Beta-site APP-cleaving
     enzyme 2) (Aspartyl protease 1) (Asp 1) (ASP1) (Membrane-associated
DE
     aspartic protease 1) (Memapsin-1) (Down region aspartic protease).
DE
GN
     BACE2 OR ASP21.
OS
     Homo sapiens (Human).
OC
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
     NCBI TaxID=9606;
OX
RN
     [1]
RP
     SEQUENCE FROM N.A.
     MEDLINE=20057170; PubMed=10591213;
RX
     Yan R., Bienkowski M.J., Shuck M.E., Miao H., Tory M.C., Pauley A.M.,
RA
     Brashier J.R., Stratman N.C., Mathews W.R., Buhl A.E., Carter D.B.,
RA
     Tomasselli A.G., Parodi L.A., Heinrikson R.L., Gurney M.E.;
RA
     "Membrane-anchored aspartyl protease with Alzheimer's disease
RT
RT
     beta-secretase activity.";
     Nature 402:533-537(1999).
RL
```

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[2]
RN
     SEQUENCE FROM N.A.
RP
RC
     TISSUE=Bone marrow;
    Xin H., Stephans J.C., Duan X., Harrowe G., Kim E., Grieshammer U.,
RA
    Giese K.;
RA
     "Identification of a novel aspartic-like protease differentially
RT
     expressed in human breast cancer cell lines.";
RT
     Submitted (JAN-1999) to the EMBL/GenBank/DDBJ databases.
RL
RN
RP
     SEQUENCE FROM N.A.
    Accarino M.P., Fumagalli P., Ottolenghi S., Taramelli R.;
RA
     "Cloning of a gene from chromosome 21 Down region encoding a potential
RT
     transmembrane aspartyl protease.";
RT
     Submitted (FEB-1998) to the EMBL/GenBank/DDBJ databases.
RL
RN
     [4]
     SEQUENCE FROM N.A.
RP
     Solans A., Estivill X., de la Luna S.;
RA
     "Cloning of a novel mammalian aspartyl protease.";
RT
     Submitted (AUG-1999) to the EMBL/GenBank/DDBJ databases.
RL
RN
     SEQUENCE FROM N.A.
RP
     MEDLINE=20120043; PubMed=10656250;
RX
     Hussain I., Powell D.J., Howlett D.R., Tew D.G., Meek T.D.,
RA
     Chapman C., Gloger I.S., Murphy K.E., Southan C.D., Ryan D.M.,
RA
     Smith T.S., Simmons D.L., Walsh F.S., Dingwall C., Christie G.;
RA
     "Identification of a novel aspartic proteinase (Asp 2) as
RT
     beta-secretase.";
RT
    Mol. Cell. Neurosci. 14:419-427(1999).
RL
RN
     SEOUENCE FROM N.A.
RP
     MEDLINE=20144060; PubMed=10677483;
RX
     Lin X., Koelsch G., Wu S., Downs D., Dashti A., Tang J.;
RA
     "Human aspartic protease memapsin 2 cleaves the beta-secretase site of
RT
     beta-amyloid precursor protein.";
RT
     Proc. Natl. Acad. Sci. U.S.A. 97:1456-1460(2000).
RL
RN
     [7]
RP
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     MEDLINE=20289799; PubMed=10830953;
RX
     Hattori M., Fujiyama A., Taylor T.D., Watanabe H., Yada T.,
RA
     Park H.-S., Toyoda A., Ishii K., Totoki Y., Choi D.-K., Groner Y.,
RA
     Soeda E., Ohki M., Takagi T., Sakaki Y., Taudien S., Blechschmidt K.,
RA
RA
     Polley A., Menzel U., Delabar J., Kumpf K., Lehmann R., Patterson D.,
     Reichwald K., Rump A., Schillhabel M., Schudy A., Zimmermann W.,
RA
     Rosenthal A., Kudoh J., Shibuya K., Kawasaki K., Asakawa S.,
RA
RA
     Shintani A., Sasaki T., Nagamine K., Mitsuyama S., Antonarakis S.E.,
     Minoshima S., Shimizu N., Nordsiek G., Hornischer K., Brandt P.,
RA
     Scharfe M., Schoen O., Desario A., Reichelt J., Kauer G., Bloecker H.,
RA
     Ramser J., Beck A., Klages S., Hennig S., Riesselmann L., Dagand E.,
RA
RA
     Wehrmeyer S., Borzym K., Gardiner K., Nizetic D., Francis F.,
     Lehrach H., Reinhardt R., Yaspo M.-L.;
RA
RΤ
     "The DNA sequence of human chromosome 21.";
RL
     Nature 405:311-319(2000).
RN
     [8]
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RP
RC
     TISSUE=Skin;
     MEDLINE=22388257; PubMed=12477932;
RX
     Strausberg R.L., Feingold E.A., Grouse L.H., Derge J.G.,
RA
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Klausner R.D., Collins F.S., Wagner L., Shenmen C.M., Schuler G.D.,
RA
    Altschul S.F., Zeeberg B., Buetow K.H., Schaefer C.F., Bhat N.K.,
RA
    Hopkins R.F., Jordan H., Moore T., Max S.I., Wang J., Hsieh F.,
RA
     Diatchenko L., Marusina K., Farmer A.A., Rubin G.M., Hong L.,
RA
     Stapleton M., Soares M.B., Bonaldo M.F., Casavant T.L., Scheetz T.E.,
RA
     Brownstein M.J., Usdin T.B., Toshiyuki S., Carninci P., Prange C.,
RA
     Raha S.S., Loquellano N.A., Peters G.J., Abramson R.D., Mullahy S.J.,
RA
     Bosak S.A., McEwan P.J., McKernan K.J., Malek J.A., Gunaratne P.H.,
RA
     Richards S., Worley K.C., Hale S., Garcia A.M., Gay L.J., Hulyk S.W.,
RA
     Villalon D.K., Muzny D.M., Sodergren E.J., Lu X., Gibbs R.A.,
RA
     Fahey J., Helton E., Ketteman M., Madan A., Rodrigues S., Sanchez A.,
RA
     Whiting M., Madan A., Young A.C., Shevchenko Y., Bouffard G.G.,
RA
     Blakesley R.W., Touchman J.W., Green E.D., Dickson M.C.,
RA
     Rodriguez A.C., Grimwood J., Schmutz J., Myers R.M.,
RA
     Butterfield Y.S.N., Krzywinski M.I., Skalska U., Smailus D.E.,
RA
     Schnerch A., Schein J.E., Jones S.J.M., Marra M.A.;
RA
     "Generation and initial analysis of more than 15,000 full-length
RT
     human and mouse cDNA sequences.";
RT
     Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903(2002).
RL
RN
     CHARACTERIZATION.
RΡ
     MEDLINE=22088158; PubMed=12093293;
RX
     Turner R.T. III, Loy J.A., Nguyen C., Devasamudram T., Ghosh A.K.,
RA
     Koelsch G., Tang J.;
RA
     "Specificity of memapsin 1 and its implications on the design of
RT
     memapsin 2 (beta-secretase) inhibitor selectivity.";
RT
     Biochemistry 41:8742-8746(2002).
RL
     -!- CATALYTIC ACTIVITY: Broad endopeptidase specificity. Cleaves Glu-
CC
         Val-Asn-Leu-|-Asp-Ala-Glu-Phe in the Swedish variant of
CC
         Alzheimer's amyloid precursor protein.
CC
     -!- SUBCELLULAR LOCATION: Type I membrane protein.
CC
     -!- SIMILARITY: Belongs to peptidase family A1.
CC
     CC
     This SWISS-PROT entry is copyright. It is produced through a collaboration
CC
     between the Swiss Institute of Bioinformatics and the EMBL outstation -
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     the European Bioinformatics Institute. There are no restrictions on its
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     use by non-profit institutions as long as its content is in no way
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     entities requires a license agreement (See http://www.isb-sib.ch/announce/
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     or send an email to license@isb-sib.ch).
CC
CC
     EMBL; AF200342; AAF17078.1; -.
DR
     EMBL; AF117892; AAD45240.1; -.
DR
     EMBL; AF050171; AAD45963.1; -.
DR
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DR
     EMBL; AF204944; AAF26368.1; -.
DR
     EMBL; AF200192; AAF13714.1; -.
DR
     EMBL; AL163284; CAB90458.1; -.
DR
     EMBL; AL163285; CAB90554.1; -.
DR
     EMBL; BC014453; AAH14453.1; -.
DR
     HSSP; P00797; 2REN.
DR
     MEROPS; A01.041; -.
DR
     Genew; HGNC:934; BACE2.
DR
DR
     MIM: 605668; -.
     GO; GO:0005624; C:membrane fraction; TAS.
DR
     GO; GO:0004190; F:aspartic-type endopeptidase activity; TAS.
DR
     GO; GO:0006464; P:protein modification; TAS.
```

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GO; GO:0009306; P:protein secretion; TAS.
DR
        InterPro; IPR001969; Aspprotease AS.
DR
        InterPro; IPR009007; Pept A acid.
DR
        InterPro; IPR001461; Peptidase Al.
DR
        Pfam; PF00026; asp; 1.
DR
        PROSITE; PS00141; ASP PROTEASE; 2.
DR
        Hydrolase; Aspartyl protease; Glycoprotein; Zymogen; Transmembrane;
KW
KW
        Signal.
                                                        POTENTIAL.
                                         20
        SIGNAL
                               1
FT
                                                        POTENTIAL.
        PROPEP
                              21
                                           ?
FT
                                        518
                                                        BETA SECRETASE 2.
        CHAIN
                               ?
FT
                                                        EXTRACELLULAR (POTENTIAL).
                              21
                                        473
FT
        DOMAIN
                                                        POTENTIAL.
                            474
                                        494
        TRANSMEM
FT
                                                        CYTOPLASMIC (POTENTIAL).
                            495
                                        518
        DOMAIN
FT
                                                        BY SIMILARITY.
        ACT SITE
                            110
                                        110
FT
        ACT SITE
                                        303
                                                        BY SIMILARITY.
                            303
FT
                                                        N-LINKED (GLCNAC. . .) (POTENTIAL).
                            170
                                        170
        CARBOHYD
FT
                                                        N-LINKED (GLCNAC. . .) (POTENTIAL).
                            366
                                        366
FΤ
        CARBOHYD
                                                        A \rightarrow T (IN REF. 6).
                              36
                                          36
        CONFLICT
FT
                                         56180 MW; 2E903150823760D3 CRC64;
                           518 AA;
        SEQUENCE
SO
                                           100.0%; Score 2687; DB 1; Length 518;
    Query Match
                                                          Pred. No. 2e-187;
                                           100.0%;
    Best Local Similarity
                                                                                                                                0;
                                                                                                            0;
                                                       0: Mismatches
                                                                                     0;
                                                                                            Indels
                                                                                                                   Gaps
    Matches 518; Conservative
                    1 MGALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGL 60
Qу
                        11. | | 11. | | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11. | 11.
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Db
                   61 ALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAG 120
Qу
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 Db
                 121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
 Qу
                        121 TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATI 180
 Db
                 181 FESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVA 240
 Qу
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                 241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
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 Qу
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 Qу
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                  421 RAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVCG 480
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 Db
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InterPro; IPR001969; Aspprotease AS.

DR

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RESULT 2
BACE RAT
                                        501 AA.
                   STANDARD;
                                 PRT;
    BACE RAT
ID
AC
    P56819;
    30-MAY-2000 (Rel. 39, Created)
DT
    30-MAY-2000 (Rel. 39, Last sequence update)
DT
    10-OCT-2003 (Rel. 42, Last annotation update)
DT
    Beta-secretase precursor (EC 3.4.23.46) (Beta-site APP cleaving
DE
    enzyme) (Beta-site amyloid precursor protein cleaving enzyme)
DE
    (Aspartyl protease 2) (Asp 2) (ASP2) (Membrane-associated aspartic
DE
    protease 2) (Memapsin-2).
DE
GN
    BACE.
    Rattus norvegicus (Rat).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Rattus.
OC
    NCBI TaxID=10116;
OX
RN
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RP
    MEDLINE=20002972; PubMed=10531052;
RX
    Vassar R., Bennett B.D., Babu-Khan S., Kahn S., Mendiaz E.A.,
RA
     Denis P., Teplow D.B., Ross S., Amarante P., Loeloff R., Luo Y.,
RA
     Fisher S., Fuller J., Edenson S., Lile J., Jarosinski M.A.,
RA
     Biere A.L., Curran E., Burgess T., Louis J.-C., Collins F.,
RA
     Treanor J., Rogers G., Citron M.;
RA
     "Beta-secretase cleavage of Alzheimer's amyloid precursor protein by
RT
     the transmembrane aspartic protease BACE.";
RT
     Science 286:735-741(1999).
RL
     -!- FUNCTION: Responsible for the proteolytic processing of the
CC
         amyloid precursor protein (APP). Cleaves at the amino terminus of
CC
         the A-beta peptide sequence, between residues 671 and 672 of APP,
CC
         leads to the generation and extracellular release of beta-cleaved
CC
         soluble APP, and a corresponding cell-associated carboxy-terminal
CC
         fragment which is later release by gamma-secretase (By
CC
         similarity).
CC
     -!- CATALYTIC ACTIVITY: Broad endopeptidase specificity. Cleaves Glu-
        Val-Asn-Leu-|-Asp-Ala-Glu-Phe in the Swedish variant of
CC
         Alzheimer's amyloid precursor protein.
CC
     -!- SUBCELLULAR LOCATION: Type I membrane protein.
CC
     -!- SIMILARITY: Belongs to peptidase family Al.
CC
     ______
CC
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CC
     between the Swiss Institute of Bioinformatics and the EMBL outstation -
CC
     the European Bioinformatics Institute. There are no restrictions on its
CC
     use by non-profit institutions as long as its content is in no way
CC
     modified and this statement is not removed. Usage by and for commercial
CC
     entities requires a license agreement (See http://www.isb-sib.ch/announce/
CC
     or send an email to license@isb-sib.ch).
CC
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CC
     EMBL; AF190727; AAF04144.1; -.
DR
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     MEROPS; A01.004; -.
DR
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        InterPro; IPR009007; Pept A acid.
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        Pfam; PF00026; asp; 1.
DR
        PRINTS; PR00792; PEPSIN.
DR
        PROSITE; PS00141; ASP PROTEASE; 1.
DR
KW
        Hydrolase; Aspartyl protease; Glycoprotein; Zymogen; Transmembrane;
KW
        Signal.
        SIGNAL
                               1
                                         21
                                                        POTENTIAL.
FT
        PROPEP
                              22
                                         45
                                                         POTENTIAL.
FT
        CHAIN
                              46
                                        501
                                                        BETA-SECRETASE.
FΤ
                              22
                                        457
                                                        EXTRACELLULAR (POTENTIAL).
        DOMAIN
FT
                            458
                                        478
                                                        POTENTIAL.
        TRANSMEM
FT
                            479
                                        501
                                                        CYTOPLASMIC (POTENTIAL).
FT
        DOMAIN
        ACT SITE
                            93
                                        93
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FT
        ACT SITE
                            289
                                        289
                                                        BY SIMILARITY.
FT
                            216
                                        420
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FT
        DISULFID
                            278
                                        443
                                                        BY SIMILARITY.
FT
        DISULFID
FT
        DISULFID
                            330
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                                                        BY SIMILARITY.
                            153
                                        153
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FT
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                                                        N-LINKED (GLCNAC. . .) (POTENTIAL).
FT
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                            172
                                        172
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FT
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                            223
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                            354
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   Best Local Similarity 46.4%; Pred. No. 1.2e-78;
   Matches 240; Conservative 82; Mismatches 165; Indels
                                                                                                           30; Gaps
                                                                                                                                 9;
                    9 LLPLLAQWLLRAAPELAPAPFT----LPLRVAAATNRVVAPTPGP--GTPAERHADGLA 61
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                        1 MAPALRWLLLWVGSGMLPAQGTHLGIRLPLRSGLA-----GPPLGLRLPRETDE-- 49
Db
                  62 LALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGT 121
Qу
                                              - 1
                  50 ---EP--EEPGRRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFAVGAA 104
Db
                122 PHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIF 181
Qv
                        105 PHPFLHRYYOROLSSTYRDLRKSVYVPYTOGKWEGELGTDLVSIPHGPNVTVRANIAAIT 164
Db
                 182 ESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPV-- 239
Qy
                        165 ESDKFFINGSNWEGILGLAYAEIARPDDSLEPFFDSLVKQTHIPNIFSLQLCGAGFPLNQ 224
Db
                 240 -AGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNAD 298
Qy
                                225 TEALASVGGSMIIGGIDHSLYTGSLWYTPIRREWYYEVIIVRVEINGQDLKMDCKEYNYD 284
Db
                 299 KAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKI 358
Qy
                        285 KSIVDSGTTNLRLPKKVFEAAVKSIKAASSTEKFPDGFWLGEQLVCWQAGTTPWNIFPVI 344
Db
                 359 SIYLRDENSSRSFRITILPQLYIQPMMGAGLNY-ECYRFGISPSTNALVIGATVMEGFYV 417
Qy
                        1:|| | :::|1|||| | | ::|: | : :||:| :| | :| | :|| :|| :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :|| | :
                 345 SLYLMGEVTNQSFRITILPQQYLRPVEDVATSQDDCYKFAVSQSSTGTVMGAVIMEGFYV 404
Db
                 418 IFDRAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMS 477
Qv
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:: | | | | | | :
                                                              | | ::| : :
              :|||:||:||
          405 VFDRARKRIGFAVSACHVHDEFRTAAVEGPFVTADMEDCGYNIPQTDESTLMTIAYVMAA 464
Db
          478 VCGAILLVLIVLLLLPFRCQR--RPRDPEVVNDESSL 512
Qу
              :| [: :: : |:: :|| | | : : :| | |
          465 IC-ALFMLPLCLMVCQWRCLRCLRHQHDDFADDISLL 500
Db
RESULT 3
BACE MOUSE
                                   PRT;
                                           501 AA.
     BACE MOUSE
                    STANDARD;
ID
AC
     P56818;
     30-MAY-2000 (Rel. 39, Created)
DT
     16-OCT-2001 (Rel. 40, Last sequence update)
DΤ
     10-OCT-2003 (Rel. 42, Last annotation update)
DT
     Beta-secretase precursor (EC 3.4.23.46) (Beta-site APP cleaving
DΕ
     enzyme) (Beta-site amyloid precursor protein cleaving enzyme)
DE
     (Aspartyl protease 2) (Asp 2) (ASP2) (Membrane-associated aspartic
DΕ
     protease 2) (Memapsin-2).
DE
     BACE.
GN
     Mus musculus (Mouse).
OS
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OC
OX
     NCBI TaxID=10090;
RN
     [1]
     SEQUENCE FROM N.A.
RP
     MEDLINE=20002972; PubMed=10531052;
RX
     Vassar R., Bennett B.D., Babu-Khan S., Kahn S., Mendiaz E.A.,
RA
     Denis P., Teplow D.B., Ross S., Amarante P., Loeloff R., Luo Y.,
RA
     Fisher S., Fuller J., Edenson S., Lile J., Jarosinski M.A.,
RA
     Biere A.L., Curran E., Burgess T., Louis J.-C., Collins F.,
RA
     Treanor J., Rogers G., Citron M.;
RA
     "Beta-secretase cleavage of Alzheimer's amyloid precursor protein by
RT
     the transmembrane aspartic protease BACE.";
RT
     Science 286:735-741(1999).
RL
RN
     [2]
RP
     REVISIONS TO 6 AND 81-87.
     Bennett B.D., Vassar R., Citron M.;
RA
     Submitted (JAN-2000) to the EMBL/GenBank/DDBJ databases.
RL
RN
     [3]
RP
     SEOUENCE FROM N.A.
     MEDLINE=20057170; PubMed=10591213;
RX
     Yan R., Bienkowski M.J., Shuck M.E., Miao H., Tory M.C., Pauley A.M.,
RA
     Brashier J.R., Stratman N.C., Mathews W.R., Buhl A.E., Carter D.B.,
RA
     Tomasselli A.G., Parodi L.A., Heinrikson R.L., Gurney M.E.;
RA
     "Membrane-anchored aspartyl protease with Alzheimer's disease
RT
     beta-secretase activity.";
RT
RL
     Nature 402:533-537(1999).
     -!- FUNCTION: Responsible for the proteolytic processing of the
CC
         amyloid precursor protein (APP). Cleaves at the amino terminus of
CC
         the A-beta peptide sequence, between residues 671 and 672 of APP,
CC
         leads to the generation and extracellular release of beta-cleaved
CC
         soluble APP, and a corresponding cell-associated carboxy-terminal
CC
         fragment which is later release by gamma-secretase (By
CC
CC
         similarity).
     -!- CATALYTIC ACTIVITY: Broad endopeptidase specificity. Cleaves Glu-
CC
         Val-Asn-Leu-|-Asp-Ala-Glu-Phe in the Swedish variant of
CC
```

```
Alzheimer's amyloid precursor protein.
CC
    -!- SUBCELLULAR LOCATION: Type I membrane protein.
CC
    -!- TISSUE SPECIFICITY: Brain.
CC
    -!- SIMILARITY: Belongs to peptidase family A1.
CC
    _____
CC
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CC
    between the Swiss Institute of Bioinformatics and the EMBL outstation -
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    the European Bioinformatics Institute. There are no restrictions on its
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    use by non-profit institutions as long as its content is in no way
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    modified and this statement is not removed. Usage by and for commercial
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    entities requires a license agreement (See http://www.isb-sib.ch/announce/
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    or send an email to license@isb-sib.ch).
CC
    ______
CC
    EMBL; AF190726; AAF04143.2; -.
DR
    EMBL; AF200346; AAF17082.1; -.
DR
    HSSP; P56272; 1AM5.
DR
    MEROPS; A01.004; -.
DR
    MGD; MGI:1346542; Bace.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR009007; Pept_A_acid.
DR
    InterPro; IPR001461; Peptidase_A1.
DR
    Pfam; PF00026; asp; 1.
DR
    PRINTS; PR00792; PEPSIN.
DR
    PROSITE; PS00141; ASP PROTEASE; 1.
DR
    Hydrolase; Aspartyl protease; Glycoprotein; Zymogen; Transmembrane;
KW
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KW
                              POTENTIAL.
    SIGNAL
                1
                      21
FT
    PROPEP
               22
                     45
                              POTENTIAL.
TЧ
    CHAIN
               46
                     501
                             BETA-SECRETASE.
FT
                             EXTRACELLULAR (POTENTIAL).
    DOMAIN
               22
                     457
FT
              458
                    478
                             POTENTIAL.
FT
   TRANSMEM
                             CYTOPLASMIC (POTENTIAL).
   DOMAIN
               479
                     501
FT
    ACT SITE
               93
                    93
                             BY SIMILARITY.
FT
    ACT SITE
               289
                    289
                             BY SIMILARITY.
FT
               216
                             BY SIMILARITY.
    DISULFID
                    420
FT
                             BY SIMILARITY.
FT
    DISULFID
               278
                    443
    DISULFID
                             BY SIMILARITY.
FT
               330
                     380
                             N-LINKED (GLCNAC. . .) (POTENTIAL).
    CARBOHYD 153 153
FΤ
                             N-LINKED (GLCNAC. . .) (POTENTIAL).
FT
    CARBOHYD 172 172
                             N-LINKED (GLCNAC. . .) (POTENTIAL).
    CARBOHYD 223 223
FT
                             N-LINKED (GLCNAC. . .) (POTENTIAL).
FT
    CARBOHYD
              354
                    354
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SO
    SEQUENCE
                       44.1%; Score 1185; DB 1; Length 501;
  Query Match
  Best Local Similarity 46.0%; Pred. No. 1.7e-78;
                                                        26; Gaps
                                                                  7;
  Matches 237; Conservative 83; Mismatches 169; Indels
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Qу
                  : 11 1
                                   1 MAPALHWLLLWVGSGMLPAQGTHLGIRLPLRSGLA-----GPPLGLRLPRETDEES---- 51
Db
          64 LEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAVAGTPH 123
Qу
                  | :|: ||||:| ||:||:|| :|:||| | ||:|||
          52 ----EEPGRRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFAVGAAPH 106
Db
         124 SYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFES 183
QV
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107 PFLHRYYQRQLSSTYRDLRKGVYVPYTQGKWEGELGTDLVSIPHGPNVTVRANIAAITES 166
Db
         184 ENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPV---A 240
Qy
             167 DKFFINGSNWEGILGLAYAEIARPDDSLEPFFDSLVKQTHIPNIFSLQLCGAGFPLNQTE 226
Db
         241 GSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKA 300
Qу
               227 ALASVGGSMIIGGIDHSLYTGSLWYTPIRREWYYEVIIVRVEINGQDLKMDCKEYNYDKS 286
Db
         301 IVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISI 360
Qу
             287 IVDSGTTNLRLPKKVFEAAVKSIKAASSTEKFPDGFWLGEQLVCWQAGTTPWNIFPVISL 346
Db
         361 YLRDENSSRSFRITILPQLYIQPMMGAGLNY-ECYRFGISPSTNALVIGATVMEGFYVIF 419
Qу
             347 YLMGEVTNQSFRITILPQQYLRPVEDVATSQDDCYKFAVSQSSTGTVMGAVIMEGFYVVF 406
Db
         420 DRAQKRVGFAASPCAEIAGAAVSEISGPFSTEDVASNCVPAQSLSEPILWIVSYALMSVC 479
QУ
                                :: ||| | |:
                                                     | | | ::| : ::|
             |||:||:||| |
         407 DRARKRIGFAVSACHVHDEFRTAAVEGPFVTADMEDCGYNIPQTDESTLMTIAYVMAAIC 466
Db
         480 GAILLVLIVLLLLPFRCQR--RPRDPEVVNDESSL 512
Qу
              |: :: : |:: :|| | | : : :| | |
         467 -ALFMLPLCLMVCQWRCLRCLRHQHDDFADDISLL 500
Db
RESULT 4
BACE HUMAN
                  STANDARD;
                                PRT;
                                      501 AA.
    BACE HUMAN
    P56817; Q9BYB9; Q9BYC0; Q9BYC1; Q9UJT5;
    30-MAY-2000 (Rel. 39, Created)
DΤ
    30-MAY-2000 (Rel. 39, Last sequence update)
DΤ
    10-OCT-2003 (Rel. 42, Last annotation update)
DT
    Beta-secretase precursor (EC 3.4.23.46) (Beta-site APP cleaving
DE.
    enzyme) (Beta-site amyloid precursor protein cleaving enzyme)
DE
     (Aspartyl protease 2) (Asp 2) (ASP2) (Membrane-associated aspartic
DE
    protease 2) (Memapsin-2).
DE
GN
    BACE OR BACE1.
OS
    Homo sapiens (Human).
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
OC
OX
    NCBI TaxID=9606;
RN
     [1]
RP
    SEQUENCE FROM N.A. (ISOFORM A).
    TISSUE=Brain;
RC.
    MEDLINE=20002972; PubMed=10531052;
RX
    Vassar R., Bennett B.D., Babu-Khan S., Kahn S., Mendiaz E.A.,
RA
     Denis P., Teplow D.B., Ross S., Amarante P., Loeloff R., Luo Y.,
RA
     Fisher S., Fuller J., Edenson S., Lile J., Jarosinski M.A.,
RA
     Biere A.L., Curran E., Burgess T., Louis J.-C., Collins F.,
RA
     Treanor J., Rogers G., Citron M.;
RA
     "Beta-secretase cleavage of Alzheimer's amyloid precursor protein by
RT
     the transmembrane aspartic protease BACE.";
RT
     Science 286:735-741(1999).
RL
RN
     SEQUENCE FROM N.A. (ISOFORM A), SEQUENCE OF 46-68, AND
RP
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RP
    CHARACTERIZATION.
RC
     TISSUE=Brain;
RX
    MEDLINE=20057171; PubMed=10591214;
     Sinha S., Anderson J.P., Barbour R., Basi G.S., Caccavello R.,
RA
     Davis D., Doan M., Dovey H.F., Frigon N., Hong J., Jacobson-Croak K.,
RA
RA
     Jewett N., Keim P., Knops J., Lieberburg I., Power M., Tan H.,
RA
     Tatsuno G., Tung J., Schenk D., Seubert P., Suomensaari S.M., Wang S.,
    Walker D., Zhao J., McConlogue L., Varghese J.;
RA
     "Purification and cloning of amyloid precursor protein beta-secretase
RT
RT
     from human brain.";
     Nature 402:537-540(1999).
RL
RN
     [3]
     SEQUENCE FROM N.A. (ISOFORM A).
RP
    MEDLINE=20057170; PubMed=10591213;
RX
     Yan R., Bienkowski M.J., Shuck M.E., Miao H., Tory M.C., Pauley A.M.,
RA
     Brashier J.R., Stratman N.C., Mathews W.R., Buhl A.E., Carter D.B.,
RA
     Tomasselli A.G., Parodi L.A., Heinrikson R.L., Gurney M.E.;
RA
     "Membrane-anchored aspartyl protease with Alzheimer's disease beta-
RT
     secretase activity.";
RТ
RL
     Nature 402:533-537(1999).
RN
     [4]
RP
     SEQUENCE FROM N.A. (ISOFORM A).
RX
    MEDLINE=20120043; PubMed=10656250;
     Hussain I., Powell D.J., Howlett D.R., Tew D.G., Meek T.D.,
RA
     Chapman C., Gloger I.S., Murphy K.E., Southan C.D., Ryan D.M.,
RA
     Smith T.S., Simmons D.L., Walsh F.S., Dingwall C., Christie G.;
RA
     "Identification of a novel aspartic proteinase (Asp 2) as beta-
RT
RT
     secretase.";
     Mol. Cell. Neurosci. 14:419-427(1999).
RL
RN
     [5]
RP
     SEQUENCE FROM N.A. (ISOFORM B).
     TISSUE=Brain, and Pancreas;
RC
     Michel B., De Pietri Tonelli D., Zacchetti D., Keller P.;
RA
     "New beta-site APP cleaving enzyme isoform (BACE-1B) obtained from
RT
     human brain and pancreas.";
RT
     Submitted (JAN-2001) to the EMBL/GenBank/DDBJ databases.
RL
RN
RP
     SEQUENCE FROM N.A. (ISOFORM C).
RC
     TISSUE=Pancreas;
     Zacchetti D., De Pietri Tonelli D., Schnurbus R.;
RA
     "New beta-site APP cleaving enzyme isoform (BACE-1C) obtained from
RT
RT
     human pancreas.";
     Submitted (JAN-2001) to the EMBL/GenBank/DDBJ databases.
RL
RN
     SEQUENCE FROM N.A. (ISOFORMS B; C AND D).
RP
RC
     TISSUE=Brain;
     MEDLINE=21408467; PubMed=11516562;
RX
RA
     Tanahashi H., Tabira T.;
     "Three novel alternatively spliced isoforms of the human beta-site
RT
     amyloid precursor protein cleaving enzyme (BACE) and their effect on
RT
     amyloid beta-peptide production.";
RT
     Neurosci. Lett. 307:9-12(2001).
RL
RN
     SEQUENCE OF 14-501 FROM N.A. (ISOFORM A), AND CHARACTERIZATION.
RP
RX
     MEDLINE=20144060; PubMed=10677483;
     Lin X., Koelsch G., Wu S., Downs D., Dashti A., Tang J.;
RA
     "Human aspartic protease memapsin 2 cleaves the beta-secretase site of
RT
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RT
    beta-amyloid precursor protein.";
RL
    Proc. Natl. Acad. Sci. U.S.A. 97:1456-1460(2000).
RN
RP
    DISULFIDE BONDS.
    MEDLINE=21950860; PubMed=11953458;
RX
    Fischer F., Molinari M., Bodendorf U., Paganetti P.;
RA
    "The disulphide bonds in the catalytic domain of BACE are critical but
RT
    not essential for amyloid precursor protein processing activity.";
RT
    J. Neurochem. 80:1079-1088(2002).
RL
    -!- FUNCTION: Responsible for the proteolytic processing of the
CC
        amyloid precursor protein (APP). Cleaves at the amino terminus of
CC
        the A-beta peptide sequence, between residues 671 and 672 of APP,
CC
        leads to the generation and extracellular release of beta-cleaved
CC
        soluble APP, and a corresponding cell-associated carboxy-terminal
CC
        fragment which is later release by gamma-secretase.
CC
    -!- CATALYTIC ACTIVITY: Broad endopeptidase specificity. Cleaves Glu-
CC
        Val-Asn-Leu-|-Asp-Ala-Glu-Phe in the Swedish variant of
CC
CC
        Alzheimer's amyloid precursor protein.
    -!- SUBCELLULAR LOCATION: Type I membrane protein.
CC
    -!- ALTERNATIVE PRODUCTS:
CC
CC
        Event=Alternative splicing; Named isoforms=4;
CC
        Name=A; Synonyms=BACE-1A, BAC-501;
CC
          IsoId=P56817-1; Sequence=Displayed;
CC
        Name=B; Synonyms=BACE-1B, BACE-I-476;
CC
          IsoId=P56817-2; Sequence=VSP 005223;
CC
        Name=C; Synonyms=BACE-1C, BACE-I-457;
          IsoId=P56817-3; Sequence=VSP 005222;
CC
CC
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CC
     -!- SIMILARITY: Belongs to peptidase family Al.
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CC
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     use by non-profit institutions as long as its content is in no way
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    modified and this statement is not removed. Usage by and for commercial
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     entities requires a license agreement (See http://www.isb-sib.ch/announce/
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     or send an email to license@isb-sib.ch).
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     EMBL; AF338816; AAK38374.1; -.
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     EMBL; AB050438; BAB40933.1; -.
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     PIR; A59090; A59090.
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     PDB; 1M4H; 28-AUG-02.
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     MEROPS; A01.004; -.
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    InterPro; IPR009007; Pept A acid.
DR
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     28-FEB-2003 (Rel. 41, Last annotation update)
DT
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     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
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     Kageyama T., Tanabe K., Koiwai O.;
     "Development-dependent expression of isozymogens of monkey
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     pepsinogens and structural differences between them.";
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     Eur. J. Biochem. 202:205-215(1991).
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RP
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    MEDLINE=86168133; PubMed=3514597;
RA
     Kageyama T., Takahashi K.;
     "The complete amino acid sequence of monkey progastricsin.";
RT
     J. Biol. Chem. 261:4406-4419(1986).
RL
RN
     [3]
RP
     SEQUENCE OF 6-65.
RX
     MEDLINE=85289106; PubMed=3928607;
RA
     Kageyama T., Takahashi K.;
     "Monkey pepsinogens and pepsins. VII. Analysis of the activation
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     process and determination of the NH2-terminal 60-residue sequence of
RT
     Japanese monkey progastricsin, and molecular evolution of
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     pepsinogens.";
     J. Biochem. 97:1235-1246(1985).
RL
CC
     -!- CATALYTIC ACTIVITY: More restricted specificity than pepsin A, but
         shows preferential cleavage at Tyr-|-Xaa bonds; high activity
CC
         towards hemoglobin as substrate.
CC
     -!- PTM: Each pepsinogen is converted to corresponding pepsin at pH
CC
         2.0 in part as a result of the release of a 47 aa activation
CC
```

```
CC
       segment and in part as a result of stepwise proteolytic cleavage
CC
       via an intermediate form(s).
    -!- MISCELLANEOUS: The expression of pepsinogen genes is regulated by
CC
       hormones and related substances.
CC
CC
    -!- SIMILARITY: Belongs to peptidase family Al.
    _____
CC
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    modified and this statement is not removed. Usage by and for commercial
    entities requires a license agreement (See http://www.isb-sib.ch/announce/
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    or send an email to license@isb-sib.ch).
CC
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DR
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    MEROPS; A01.003; -.
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Qу
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Db
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Qу
                              :|||:||:||:||:||
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    Hayano T., Sogawa K., Ichihara Y., Fujii-Kuriyama Y., Takahashi K.;
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     "Primary structure of human pepsinogen C gene.";
RT
     J. Biol. Chem. 263:1382-1385(1988).
RL
RN
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     Taggart R.T., Cass L.G., Mohandas T.K., Derby P., Barr P.J., Pals G.,
RA
RA
     Bell G.I.;
     "Human pepsinogen C (progastricsin). Isolation of cDNA clones,
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     localization to chromosome 6, and sequence homology with pepsinogen
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     Samloff I.M., Walz D.A., Barr P.J., Taggart R.T.;
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     Genomics 4:137-148(1989).
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     Wong R.N.S., Tang J.;
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    Athauda S.B.P., Tanji M., Kageyama T., Takahashi K.;
     "A comparative study on the NH2-terminal amino acid sequences and
RT
     some other properties of six isozymic forms of human pepsinogens and
RT
RT
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     J. Biochem. 106:920-927(1989).
RL
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    SEQUENCE OF 17-64.
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    MEDLINE=83079318; PubMed=6816595;
RX
     Foltmann B., Jensen A.L.;
RA
     "Human progastricsin. Analysis of intermediates during activation
RT
     into gastricsin and determination of the amino acid sequence of the
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RT
     Eur. J. Biochem. 128:63-70(1982).
RL
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    X-RAY CRYSTALLOGRAPHY (1.62 ANGSTROMS).
RP
    MEDLINE=95230687; PubMed=7714902;
RX
    Moore S.A., Sielecki A.R., Chernaia M.M., Tarasova N.I., James M.N.G.;
RA
     "Crystal and molecular structures of human progastricsin at 1.62-A
RТ
RT
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     J. Mol. Biol. 247:466-485(1995).
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    MEDLINE=98069649; PubMed=9406551;
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     Khan A.R., Cherney M.M., Tarasova N.I., James M.N.;
     "Structural characterization of activation 'intermediate 2' on the
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     -!- CATALYTIC ACTIVITY: More restricted specificity than pepsin A, but
CC
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CC
         towards hemoglobin as substrate.
     -!- SIMILARITY: Belongs to peptidase family A1.
CC
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CC
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     the European Bioinformatics Institute. There are no restrictions on its
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     use by non-profit institutions as long as its content is in no way
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     or send an email to license@isb-sib.ch).
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     InterPro; IPR009007; Pept A acid.
DR
     InterPro; IPR001461; Peptidase Al.
DR
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DR
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FΤ
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            11:: 11:: 11
                                            |: |: ||||| :| |
          50 PAWKYRFGDLSVTYEP-----MAYMD-----AAYFGEISIGTPPQNFLVLFD 91
        111 TGSSNFAV-----AGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDL 162
Qу
                          92 TGSSNLWVPSVYCQSQACTSHS----RFNPSESSTYSTNGQTFSLQYGSGSLTGFFGYDT 147
Db
         163 VTIPKGFNTSFLVNIATIFESENFFLPG----IKWNGILGLAYATLAKPSSSLETFFDS 217
Qу
            148 LTV----OSIQVPNOEFGLSEN--EPGTNFVYAQFDGIMGLAYPALSVDEAT--TAMQG 198
Db
         218 LVTQANIPN-VFSMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQI 276
Qу
```

```
Db
         199 MVQEGALTSPVFSVYLSNQ-----QGSSGGAVVFGGVDSSLYTGQIYWAPVTQELYWQI 252
         277 EILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGF 336
Qу
              | : | | | | : | | : | : | : | : | : : |
         253 GIEEFLIGGQASGW-CSE--GCQAIVDTGTSLLTVPQQYMSALLQA----- 295
Db
         337 WTGSQLACWTNSETPWSYF-----PKISIYLRDENSSRSFRITILPQLYIQPMMG 386
Qу
              Db
         296 -TGAQ------EDEYGQFLVNCNSIQNLPSLTFII-----NGVEFPLPPSSYI---- 336
         387 AGLNYECY-RFGISP----STNA---LVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qу
              |: | |: | | | ::| ::|
Db
         337 --LSNNGYCTVGVEPTYLSSQNGQPLWILGDVFLRSYYSVYDLGNNRVGFATA 387
RESULT 7
PEPC CALJA
    PEPC CALJA
                STANDARD;
                              PRT; 388 AA.
AC
    Q9N2D3;
    16-OCT-2001 (Rel. 40, Created)
DТ
    16-OCT-2001 (Rel. 40, Last sequence update)
DT
DТ
    10-OCT-2003 (Rel. 42, Last annotation update)
DE
    Gastricsin precursor (EC 3.4.23.3) (Pepsinogen C).
GN
OS
    Callithrix jacchus (Common marmoset).
OC
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
    Mammalia; Eutheria; Primates; Platyrrhini; Callitrichidae;
OC
OC
    Callithrix.
    NCBI_TaxID=9483;
OX
RN
    [1]
    SEQUENCE FROM N.A., SEQUENCE OF 17-26, FUNCTION, AND ENZYME
RP
RP
    REGULATION.
RC
    TISSUE=Gastric mucosa;
RX
    MEDLINE=20250834; PubMed=10788784;
RA
    Kaqeyama T.;
RT
    "New World monkey pepsinogens A and C, and prochymosins. Purification,
RT
    characterization of enzymatic properties, cDNA cloning, and molecular
RT
    evolution.";
RL
    J. Biochem. 127:761-770(2000).
CC
    -!- FUNCTION: Hydrolyzes a variety of proteins.
    -!- CATALYTIC ACTIVITY: More restricted specificity than pepsin A, but
CC
CC
        shows preferential cleavage at Tyr-|-Xaa bonds; high activity
CC
        towards hemoglobin as substrate.
CC
    -!- ENZYME REGULATION: Inhibited by pepstatin.
CC
    -!- MISCELLANEOUS: The optimal pH is around 2.
    -!- SIMILARITY: Belongs to peptidase family Al.
CC
CC
    _____
CC
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    between the Swiss Institute of Bioinformatics and the EMBL outstation -
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    modified and this statement is not removed. Usage by and for commercial
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CC
CC
    or send an email to license@isb-sib.ch).
    _____
CC
    EMBL; AB038385; BAA90872.1; -.
DR
DR
    PIR; JC7246; JC7246.
```

```
HSSP; P20142; 1AVF.
DR
DR
    MEROPS; A01.003; -.
DR
    InterPro; IPR001969; Aspprotease AS.
    InterPro; IPR009007; Pept A acid.
DR
    InterPro; IPR001461; Peptidase Al.
DR
DR
    Pfam; PF00026; asp; 1.
    PRINTS; PR00792; PEPSIN.
DR
   PROSITE; PS00141; ASP PROTEASE; 2.
DR
    Hydrolase; Aspartyl protease; Zymogen; Digestion; Signal.
KW
FT
    SIGNAL
              1
                    16
   PROPEP
              17
                    59
                            ACTIVATION PEPTIDE (BY SIMILARITY).
FT
   CHAIN
              60
                    388
                            GASTRICSIN.
FT
            91 91
276 276
   ACT SITE
                   91
                            BY SIMILARITY.
FT
    ACT SITE
FT
                            BY SIMILARITY.
   DISULFID 104
FT
                   109
                            BY SIMILARITY.
   DISULFID 267
                    271
                            BY SIMILARITY.
FT
                   343
FT
   DISULFID 310
                           BY SIMILARITY.
SQ
    SEQUENCE 388 AA; 42503 MW; 0BC48DBD1F7D2D8C CRC64;
 Query Match 13.1%; Score 351.5; DB 1; Length 388; Best Local Similarity 30.1%; Pred. No. 3.5e-18;
 Matches 112; Conservative 56; Mismatches 115; Indels 89; Gaps
                                                                17;
         92 YYLEMLIGTPPOKLQILVDTGSSNFAV-----AGTPHSYIDTYFDTERSSTYRSKGF 143
Qу
           73 YFGEISIGTPPQNFLVLFDTGSSNLWVPSVYCQSQACTSHS----RFNPSASSTYSSNGQ 128
Db
        144 DVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFESENFFLPG----IKWNGILG 198
Qу
             129 TFSLQYGSGSLTGFFGYDTLTV----QSIQVPNQEFGLSEN--EPGTNFVYAQFDGIMG 181
Db
        199 LAYATLAKPSSSLETFFDSLVTQANIPN-VFSMQMCGAGLPVAGSGTNGGSLVLGGIEPS 257
Qу
           182 LAYPALSMGGAT--TAMQGMLQEGALTSPVFSFYLSNQ-----QGSSGGAVIFGGVDSS 233
Db
        258 LYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFD 317
Qу
           234 LYTGQIYWAPVTQELYWQIGIEEFLIGGQASGW-CSE--GCQAIVDTGTSLLTVPQQYMS 290
Db
        318 AVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYF-----PKISIYLRDENS 367
Qу
           | :: :
        291 AFLEA-----TGAO-----EDEYGOFLVNCDSIONLPTLTFII---- 323
Db
        368 SRSFRITILPOLYIOPMMGAGLNYECY-RFGISP----STNALVIGATVMEGFYVIF 419
Qy
                 : | | | : | : | : | : | : | : |
        324 -NGVEFPLPPSSYI-----LSNNGYCTVGVEPTYLSSQNSQPLWILGDVFLRSYYSVF 375
Db
        420 DRAOKRVGFAAS 431
Qу
           - 1
               | | | | | | :
Db
        376 DLGNNRVGFATA 387
RESULT 8
PEPC CAVPO
ID PEPC_CAVPO
               STANDARD; PRT; 394 AA.
AC
   064411;
    15-JUL-1998 (Rel. 36, Created)
DT
```

```
DΤ
     15-JUL-1998 (Rel. 36, Last sequence update)
DT
     28-FEB-2003 (Rel. 41, Last annotation update)
DE
     Gastricsin precursor (EC 3.4.23.3) (Pepsinogen C).
GN
OS
     Cavia porcellus (Guinea pig).
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC.
OC
    Mammalia; Eutheria; Rodentia; Hystricognathi; Caviidae; Cavia.
OX
    NCBI TaxID=10141;
RN
     [1]
RP
     SEQUENCE FROM N.A.
    MEDLINE=92355614; PubMed=1644829;
RX
     Kageyama T., Ichinose M., Tsukada S., Miki K., Kurokawa K., Koiwai O.,
RA
     Tanji M., Yakabe E., Athauda S.B., Takahashi K.;
RA
     "Gastric procathepsin E and progastricsin from quinea pig.
RT
     Purification, molecular cloning of cDNAs, and characterization of
RT
RT
     enzymatic properties, with special reference to procathepsin E.";
RL
     J. Biol. Chem. 267:16450-16459(1992).
     -!- CATALYTIC ACTIVITY: More restricted specificity than pepsin A, but
CC
CC
        shows preferential cleavage at Tyr-|-Xaa bonds; high activity
CC
        towards hemoglobin as substrate.
CC
     -!- SIMILARITY: Belongs to peptidase family A1.
     ______
CC
CC
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    use by non-profit institutions as long as its content is in no way
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    modified and this statement is not removed. Usage by and for commercial
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CC
    or send an email to license@isb-sib.ch).
CC
    EMBL; M88652; AAA37053.1; -.
DR
    PIR; B43356; B43356.
DR
DR
    HSSP; P20142; 1AVF.
DR
    MEROPS; A01.003; -.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR009007; Pept A acid.
DR
    InterPro; IPR001461; Peptidase A1.
DR
    Pfam; PF00026; asp; 1.
    PRINTS; PR00792; PEPSIN.
DR
    PROSITE; PS00141; ASP PROTEASE; 2.
DR
KW
    Hydrolase; Aspartyl protease; Zymogen; Digestion; Signal.
FT
    SIGNAL
                       16
                                POTENTIAL.
                 1
                 17
    PROPEP
                       65
FT
                                ACTIVATION PEPTIDE.
    CHAIN
                 66
                       394
FΤ
                                GASTRICSIN.
    ACT SITE
                97
                       97
                                BY SIMILARITY.
FT
    ACT SITE
                283
                       283
                                BY SIMILARITY.
FT
                                BY SIMILARITY.
FT
    DISULFID
                110
                       115
FT
    DISULFID
                273
                       277
                                BY SIMILARITY.
FT
    DISULFID
                316
                       349
                                BY SIMILARITY.
               394 AA; 42995 MW; 114F08E105D49865 CRC64;
SO
    SEQUENCE
 Query Match
                         12.1%; Score 324.5; DB 1; Length 394;
 Best Local Similarity
                         29.0%; Pred. No. 3.2e-16;
 Matches 107; Conservative 63; Mismatches 116; Indels
                                                              83;
                                                                         18;
          92 YYLEMLIGTPPQKLQILVDTGSSNF-----AVAGTPHSYIDTYFDTERSSTYRSKGF 143
Qу
             1: :: :||||| |:| ||||||
```

```
79 YFGQISLGTPPQSFQVLFDTGSSNLWVPSVYCSSLACTTH----TRFNPRDSSTYVATDQ 134
Db
         144 DVTVKYTQGSWTGFVGEDLVTI----PK-GFNTSFLVNIATIFESENFFLPG----IK 192
Qy
               1:1
                                                           - 11
         135 SFSLEYGTGSLTGVFGYDTMTIQDIQVPKQEFGLS-----ETE----PGSDFVYAE 181
Db
         193 WNGILGLAYATLAKPSSSLETFFDSLVTQANI-PNVFSMQMCGAGLPVAGS--GTNGGSL 249
Qy
                                                         | | | | | | | |
             ::||||| | |:: :: | | |:: : ::||::
         182 FDGILGLGYPGLSEGGAT--TAMQGLLREGALSQSLFSVYL------GSQQGSDEGQL 231
Db
         250 VLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLL 309
Qy
             232 ILGGVDESLYTGDIYWTPVTQELYWQIGIEGFLIDGSASGWCSR---GCQGIVDTGTSLL 288
Db
         310 RLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSR 369
Qy
                    :|:|: |: :| ::| : : | | :
             : |
         289 TVPSDYLSTLVQAIGAEE--NEYGEYF-----VSCSSIQDLPTLTFVISGV------ 332
Db
         370 SFRITILPQLYIQP----MMGAGLNYECYRFGISPSTN--ALVIGATVMEGFYVIFDRA 422
Qу
                 : | ||
                              Db
         333 --EFPLSPSAYILSGENYCMVGLESTY-----VSPGGGEPVWILGDVFLRSYYSVYDLA 384
         423 QKRVGFAAS 431
Qу
               11111:
Db
         385 NNRVGFATA 393
RESULT 9
RENI MOUSE
                            PRT;
                                       402 AA.
    RENI MOUSE
                 STANDARD;
ΤD
    P06281; P97911; Q62153; Q62154;
AC
    01-JAN-1988 (Rel. 06, Created)
DT
    01-JAN-1988 (Rel. 06, Last sequence update)
DΤ
    15-MAR-2004 (Rel. 43, Last annotation update)
DT
    Renin 1 precursor (EC 3.4.23.15) (Angiotensinogenase) (Kidney renin).
DF.
    REN1 OR REN-1 OR REN.
GN
    Mus musculus (Mouse).
OS
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
    Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Mus.
OC
    NCBI TaxID=10090;
OX
RN
    SEQUENCE FROM N.A.
RP
RC
    STRAIN=BALB/c;
    MEDLINE=84182525; PubMed=6370686;
RX
    Holm I., Ollo R., Panthier J.-J., Rougeon F.;
RA
     "Evolution of aspartyl proteases by gene duplication: the mouse renin
RT
    gene is organized in two homologous clusters of four exons.";
RT
     EMBO J. 3:557-562(1984).
RL
RN
     [2]
     SEOUENCE FROM N.A.
RP
     STRAIN=BALB/c; TISSUE=Kidney;
RC
    MEDLINE=90067953; PubMed=2685761;
RX
     Kim W.S., Murakami K., Nakayama K.;
RA
     "Nucleotide sequence of a cDNA coding for mouse Ren1 preprorenin.";
RТ
    Nucleic Acids Res. 17:9480-9480(1989).
RL
RN
     SEQUENCE FROM N.A.
RP
```

```
RC
     STRAIN=DBA/2, and C57BL/10;
    MEDLINE=90108722; PubMed=2691339;
RX
     Burt D.W., Mullins L.J., George H., Smith G., Brooks J., Pioli D.,
RA
RA
     Brammar W.J.;
RT
     "The nucleotide sequence of a mouse renin-encoding gene, Ren-1d, and
RT
     its upstream region.";
RL
    Gene 84:91-104(1989).
RN
RP
     SEQUENCE OF 1-30 FROM N.A.
RC
    TISSUE=Kidney;
RX
    MEDLINE=84298161; PubMed=6089205;
     Panthier J.-J., Dreyfus M., Roux D.T.L., Rougeon F.;
RA
RT
     "Mouse kidney and submaxillary gland renin genes differ in their 5'
RТ
     putative regulatory sequences.";
     Proc. Natl. Acad. Sci. U.S.A. 81:5489-5493(1984).
RL
RN
     SEQUENCE OF 1-31 FROM N.A.
RP
RX
    MEDLINE=85085936; PubMed=6392850;
     Field L.J., Philbrick W.M., Howles P.N., Dickinson D.P.,
RA
RA
     McGowan R.A., Gross K.W.;
RT
     "Expression of tissue-specific Ren-1 and Ren-2 genes of mice:
RT
     comparative analysis of 5'-proximal flanking regions.";
RL
    Mol. Cell. Biol. 4:2321-2331(1984).
RN
RP
     SEQUENCE OF 22-37 AND 72-80.
     STRAIN=C57BL/10ROS X C3H/HEROS; TISSUE=Kidney;
RC
     MEDLINE=97182599; PubMed=9030738;
RX
     Jones C.A., Petrovic N., Novak E.K., Swank R.T., Sigmund C.D.,
RA
RA
     Gross K.W.;
     "Biosynthesis of renin in mouse kidney tumor As4.1 cells.";
RT
     Eur. J. Biochem. 243:181-190(1997).
RL
     -!- FUNCTION: Renin is a highly specific endopeptidase, whose only
CC
         known function is to generate angiotensin I from angiotensinogen
CC
CC
         in the plasma, initiating a cascade of reactions that produce an
         elevation of blood pressure and increased sodium retention by the
CC
CC
         kidney.
     -!- CATALYTIC ACTIVITY: Cleaves Leu-|- bond in angiotensinogen to
CC
CC
         generate angiotensin I.
     -!- SUBCELLULAR LOCATION: Secreted.
CC
     -!- TISSUE SPECIFICITY: Kidney.
CC
     -!- INDUCTION: Renal renin is synthesized by the juxtaglomerular cells
CC
         of the kidney in response to decreased blood pressure and sodium
CC
CC
         concentration.
     -!- POLYMORPHISM: In inbred mouse strains, there are at least two
CC
         alleles which can occur at the Renl locus: Ren-1D and Ren-1C.
CC
CC
         The sequence shown is that of Ren-1C.
     -!- SIMILARITY: Belongs to peptidase family A1.
CC
     ______
CC
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     between the Swiss Institute of Bioinformatics and the EMBL outstation -
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     the European Bioinformatics Institute. There are no restrictions on its
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     modified and this statement is not removed. Usage by and for commercial
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CC
     or send an email to license@isb-sib.ch).
CC
     EMBL; X00810; CAA25391.1; -.
DR
```

```
DR
     EMBL; X00811; CAA25391.1; JOINED.
DR
     EMBL; X00812; CAA25391.1; JOINED.
     EMBL; X00813; CAA25391.1; JOINED.
DR
     EMBL; X00814; CAA25391.1; JOINED.
DR
     EMBL; X00815; CAA25391.1; JOINED.
DR
DR
     EMBL; X00816; CAA25391.1; JOINED.
DR
     EMBL; X00850; CAA25391.1; JOINED.
DR
     EMBL; X00851; CAA25391.1; JOINED.
     EMBL; X16642; CAA34636.1; -.
DR
DR
     EMBL; K02596; AAA40045.1; -.
     EMBL; M32352; AAA40043.1; -.
DR
     EMBL; K02800; AAA40044.1; -.
DR
DR
     EMBL; M34190; AAA40042.1; -.
DR
     PIR; A00989; REMSK.
     HSSP; P00796; 1SMR.
DR
     MEROPS; A01.007; -.
DR
     MGD; MGI:97898; Ren1.
DR
DR
     InterPro; IPR001969; Aspprotease AS.
DR
     InterPro; IPR009007; Pept A acid.
DR
     InterPro; IPR001461; Peptidase A1.
DR
     Pfam; PF00026; asp; 1.
DR
     PRINTS; PR00792; PEPSIN.
DR
     PROSITE; PS00141; ASP PROTEASE; 2.
     Hydrolase; Aspartyl protease; Plasma; Glycoprotein; Zymogen;
KW
KW
     Signal.
     SIGNAL
                          21
FT
                                   ACTIVATION PEPTIDE.
     PROPEP
                  22
                          71
FT
                  72
                         402
     CHAIN
                                   RENIN 1.
FT
     ACT SITE
                 102
                         102
                                   BY SIMILARITY.
FT
     ACT SITE
FT
                 287
                         287
                                   BY SIMILARITY.
                                   BY SIMILARITY.
FT
     DISULFID
                 115
                         122
                 278
                         282
                                   BY SIMILARITY.
FT
     DISULFID
                                   N-LINKED (GLCNAC. . .) (POTENTIAL).
                  69
                         69
FT
     CARBOHYD
                                   N-LINKED (GLCNAC. . .) (POTENTIAL).
                 139
                         139
FΤ
     CARBOHYD
                  320
                         320
                                   N-LINKED (GLCNAC. . .) (POTENTIAL).
FT
     CARBOHYD
                  58
                         58
                                   W \rightarrow R \text{ (in Ren-1D)}.
     VARIANT
FT
                                   T \rightarrow I \text{ (in Ren-1D)}.
                         68
     VARIANT
                  68
FT
                                   S \rightarrow V \text{ (in Ren-1D)}.
FT
     VARIANT
                  160
                         160
                  315
                         315
                                   E \rightarrow D (in Ren-1D).
FT
     VARIANT
FT
                  352
                         352
                                   N \rightarrow Y \text{ (in Ren-1D)}.
     VARIANT
                   6
                          23
                                   MISSING (IN REF. 1).
FT
     CONFLICT
                  24
                                   T \rightarrow I (IN REF. 1).
FT
     CONFLICT
                          24
                 163
                         163
                                   V \rightarrow VSRV (IN REF. 1).
FΤ
     CONFLICT
SO
                 402 AA; 44342 MW; D42920B555E97A38 CRC64;
     SEQUENCE
                           11.9%; Score 320; DB 1; Length 402;
  Query Match
  Best Local Similarity 28.6%; Pred. No. 7e-16;
                                66; Mismatches 181; Indels
                                                                   68; Gaps
  Matches 126; Conservative
           10 LPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPG-PGTPAERHADGLALALE--- 65
Qу
               : | | | | | |
                             : | |: || | |
                                                : |
                                                              11 1
            6 MPLWALLLL-----WSPCTFSLPTRTATFERIPLKKMPSVREILEERGVDMTRLSAEWGV 60
Db
           66 ----PA---LASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSNFAV 118
Qу
                                1:1
                                     1:
                        \perp
           61 FTKRPSLTNLTSPVVLTNYL----NTQ-----YYGEIGIGTPPQTFKVIFDTGSANLWV 110
Db
```

```
119 AGTPHSY-----IDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTS 172
Qу
                         | : ::: ||:|
                                        111 PSTKCSRLYLACGIHSLYESSDSSSYMENGSDFTIHYGSGRVKGFLSQDSVTV-GGITVT 169
Db
         173 FLVNIATIFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANI-PNVFSMQ 231
QУ
                          | |
                              |::|:||: : |:
                                                 :
                                                     | | | :::| : | | | ::
         170 QTFGEVTELPLIPFML--AKFDGVLGMGFP--AQAVGGVTPVFDHILSQGVLKEEVFSVY 225
Db
         232 MCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLD 291
QУ
                           11
         226 Y-----NRGSHLLGGEVVLGGSDPQHYQGNFHYVSISKTDSWQITMKGVSVG--SSTLL 277
Db
         292 CREYNADKAIVDSGTTLLRLPQKVFDAVVEAV-ARASLIPEFSDGFWTGSQLACWTNSET 350
Qу
                    :||:|:: : |
                                       :::|: |:
                                                11:
             278 CEEGCA--VVVDTGSSFISAPTSSLKLIMQALGAKEKRIEEY-----VVNC---SQV 324
Db
         351 PWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGL-NYECYRFGISPSTNAL-VIG 408
Qу
                              |:: :: : |
                                                 - 1
                                                       :
                                                            1 1 1
                 1 11 1
         325 P--TLPDISFDL----GGRAYTLSSTDYVLQYPNRRDKLCTLALHAMDIPPPTGPVWVLG 378
Db
         409 ATVMEGFYVIFDRAQKRVGFA 429
Qу
             11: 11
                           1:11
         379 ATFIRKFYTEFDRHNNRIGFA 399
Db
RESULT 10
CATD CLUHA
ID
     CATD CLUHA
                   STANDARD;
                                  PRT;
                                        396 AA.
AC
     Q9DEX3;
DT
     28-FEB-2003 (Rel. 41, Created)
     28-FEB-2003 (Rel. 41, Last sequence update)
DТ
     10-OCT-2003 (Rel. 42, Last annotation update)
DT
DE
     Cathepsin D precursor (EC 3.4.23.5).
OS
     Clupea harengus (Atlantic herring).
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Actinopterygii; Neopterygii; Teleostei; Clupeomorpha; Clupeidae;
OC
OC
     Clupea.
     NCBI TaxID=7950;
OX
RN
     [1]
RP
     SEQUENCE FROM N.A.
     Nielsen L.B., Stougaard P., Andersen P.S., Pedersen L.H.;
RA
     "Cloning and sequence determination of herring muscle cathepsin D.";
RT
     Submitted (OCT-2000) to the EMBL/GenBank/DDBJ databases.
RL
RN
     [2]
RP
     SEQUENCE OF 62-82.
RC-
     TISSUE=Skeletal muscle;
     MEDLINE=21165469; PubMed=11207447;
RX
     Nielsen L.B., Nielsen H.H.;
RA
     "Purification and characterization of cathepsin D from herring muscle
RT
RT
     (Clupea harengus).";
     Comp. Biochem. Physiol. 128B:351-363(2001).
RL
     -!- FUNCTION: Cathepsin D is an acid protease active in intracellular
CC
         protein breakdown.
CC
     -!- CATALYTIC ACTIVITY: Specificity similar to, but narrower than,
CC
         that of pepsin A. Does not cleave the 4-Gln-|-His-5 bond in B
CC
CC
         chain of insulin.
     -!- ENZYME REGULATION: Inhibited by pepstatin.
CC
```

```
-!- SUBUNIT: Monomer.
CC
    -!- SUBCELLULAR LOCATION: Lysosomal.
CC
    -!- MISCELLANEOUS: The isoelectric point is 6.8. Has optimal activity
CC
       at pH 2.5 with hemoglobin as the substrate and the optimal
CC
       temperature is 37 degrees Celsius.
CC
    -!- SIMILARITY: Belongs to peptidase family A1.
CC
    _____
CC
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    the European Bioinformatics Institute. There are no restrictions on its
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    use by non-profit institutions as long as its content is in no way
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    modified and this statement is not removed. Usage by and for commercial
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    entities requires a license agreement (See http://www.isb-sib.ch/announce/
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    or send an email to license@isb-sib.ch).
CC
    ______
CC
    EMBL; AF312364; AAG27733.1; -.
DR
    HSSP; P07339; 1LYB.
DR
    MEROPS; A01.009; -.
DR
    InterPro; IPR001969; Aspprotease AS.
DR
    InterPro; IPR009007; Pept_A_acid.
DR
    InterPro; IPR001461; Peptidase_A1.
DR
    Pfam; PF00026; asp; 1.
DR
    PRINTS; PR00792; PEPSIN.
DR
    PROSITE; PS00141; ASP PROTEASE; 2.
DR
    Hydrolase; Aspartyl protease; Glycoprotein; Lysosome; Signal; Zymogen.
KW
                    18
61
                             POTENTIAL.
               1
FT
    SIGNAL
                             ACTIVATION PEPTIDE.
               19
FT
    PROPEP
    CHAIN
               62
                     396
                             CATHEPSIN D.
FT
    ACT_SITE 94 94
ACT_SITE 281 281
DISULFID 107 114
                    94
                             BY SIMILARITY.
FT
                            BY SIMILARITY.
FT
                            BY SIMILARITY.
FT
                            BY SIMILARITY.
    DISULFID 272 276
FT
                           BY SIMILARITY.
N-LINKED (GLCNAC. . .) (POTENTIAL).
    DISULFID 315 352
FT
    CARBOHYD
              131
                    131
FT
    SEQUENCE 396 AA; 43315 MW; D0375DC38567A31B CRC64;
SO
 Query Match 11.7%; Score 315.5; DB 1; Length 396; Best Local Similarity 27.1%; Pred. No. 1.5e-15;
 Matches 112; Conservative 65; Mismatches 141; Indels
         50 GTPAERHADGLALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILV 109
Qу
            47 GTNSLQHNQGFPSSNAP---TPETLKNYM-----DAQYYGEIGLGTPVQMFTVVF 93
Db
        110 DTGSSNFAVAGTPHSYIDT-----YFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLV 163
Qу
            94 DTGSSNLWLPSIHCSFTDIACLLHHKYNGAKSSTYVKNGTEFAIQYGSGSLSGYLSQDSC 153
Db
         164 TIPKGFNTSFLVNIATIFESENFFLPGI----KWNGILGLAYATLAKPSSSLETFFDSL 218
Qу
            154 TI-----GDIVVEKQLF-GEAIKQPGVAFIAAKFDGILGMAYPRIS--VDGVPPVFDMM 204
Db
        219 VTQANI-PNVFSMQMCGAGLPVAGSGTN----GGSLVLGGIEPSLYKGDIWYTPIKEEW 272
Qy
            205 MSQKKVEQNVFSFYL-----NRNPDTEPGGELLLGGTDPKYYTGDFNYVPVTRQA 254
         273 YYOIEILKLEIGGOSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLI--- 329
Qy
```

```
1:|| : : || | | | | | :: :||||:||:|: |
                                                      |: :|:
         255 YWQIHMDGMSIGSQ-LTL-CKD--GCEAIVDTGTSLITGPPAEVRALQKAIGAIPLIQGE 310
Db
         330 -----PEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITILPQ 378
Qy
                       | | | | : | | | | | :: :: :: |
         311 YMIDCKKVPTLPTIS--FNVGGK----TYSLTGEQY-----VLKESQGGKTICLSGLMG 358
Db
         379 LYIQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAAS 431
Qу
                       |: ::| : :| :||
         359 LEIPP-----PAGPLWILGDVFIGQYYTVFDRESNRVGFAKS 395
Db
RESULT 11
APR1 ORYSA
                                       509 AA.
    APR1 ORYSA
                  STANDARD;
                                 PRT:
AC
    042456;
    01-NOV-1997 (Rel. 35, Created)
DT
    01-NOV-1997 (Rel. 35, Last sequence update)
DT
    28-FEB-2003 (Rel. 41, Last annotation update)
DT
    Aspartic proteinase oryzasin 1 precursor (EC 3.4.23.-).
DE
    Oryza sativa (Rice).
OS
    Eukaryota; Viridiplantae; Streptophyta; Embryophyta; Tracheophyta;
OC
    Spermatophyta; Magnoliophyta; Liliopsida; Poales; Poaceae;
OC
    Ehrhartoideae; Oryzeae; Oryza.
OC
OX
    NCBI TaxID=4530;
RN
    [1]
    SEQUENCE FROM N.A.
RP
    STRAIN=cv. Nipponbare / Japonica; TISSUE=Seed;
RC
RX
    MEDLINE=96048031; PubMed=7556174;
    Asakura T., Watanabe H., Abe K., Arai S.;
RA
    "Rice aspartic proteinase, oryzasin, expressed during seed ripening
RT
    and germination, has a gene organization distinct from those of
RT
    animal and microbial aspartic proteinases.";
RT
    Eur. J. Biochem. 232:77-83(1995).
RL
    -!- DEVELOPMENTAL STAGE: Seed ripening and germination.
CC
     -!- SIMILARITY: Belongs to peptidase family A1.
CC
     ______
CC
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     or send an email to license@isb-sib.ch).
CC
     ______
CC
DR
     EMBL; D32165; BAA06876.1; -.
     EMBL; D32144; BAA06875.1; -.
DR
DR
     PIR; S66516; S66516.
     HSSP; P42210; 1QDM.
DR
DR
    MEROPS; A01.020; -.
DR
     Gramene; Q42456; -.
     InterPro; IPR001969; Aspprotease AS.
DŔ
     InterPro; IPR009007; Pept_A_acid.
DR
     InterPro; IPR001461; Peptidase A1.
DR
     InterPro; IPR007856; SapB 1.
DR
     InterPro; IPR008138; SapB 2.
     InterPro; IPR008140; SapB sub.
DR
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InterPro; IPR008373; Saposin.
DR
    InterPro; IPR008139; SaposinB.
DR
    Pfam; PF00026; asp; 1.
DR
    Pfam; PF05184; SapB 1; 1.
DR
    Pfam; PF03489; SapB 2; 1.
DR
    PRINTS; PR00792; PEPSIN.
DR
    PRINTS; PR01797; SAPOSIN.
DR
    ProDom; PD001732; SapB sub; 1.
DR
    SMART; SM00118; SAPB; 2.
DR
    PROSITE; PS00141; ASP PROTEASE; 2.
DR
    Hydrolase; Aspartyl protease; Zymogen; Glycoprotein; Signal.
KW
                              POTENTIAL.
                1
                      20
    SIGNAL
FT
                      67
                              POTENTIAL.
                21
    PROPEP
FT
                              ASPARTIC PROTEINASE ORYZASIN 1.
                     509
                68
    CHAIN
FT
                              SPECIFIC TO PLANT ASPARTIC PROTEINASES
               318
                     416
FT
    DOMAIN
                              (BY SIMILARITY).
FT
               103
                              BY SIMILARITY.
FT
                     103
    ACT SITE
    ACT SITE
                     290
                              BY SIMILARITY.
               290
FT
                              BY SIMILARITY.
    DISULFID
               116
                     122
FT
                     285
                              BY SIMILARITY.
    DISULFID
               281
FT
               428
                     465
                              BY SIMILARITY.
    DISULFID
FΤ
                              N-LINKED (GLCNAC. . .) (POTENTIAL).
               252
                     252
FT
    CARBOHYD
                              N-LINKED (GLCNAC. . .) (POTENTIAL).
                     400
    CARBOHYD
               400
FT
              509 AA; 54145 MW; 182F5DADA4CBE358 CRC64;
    SEQUENCE
SO
                       11.7%; Score 313.5; DB 1; Length 509;
  Query Match
  Best Local Similarity 23.0%; Pred. No. 2.9e-15;
 Matches 127; Conservative 75; Mismatches 179; Indels 171; Gaps
           3 ALARALLLPLLAQWLLRAAPELAPAPFTLPLRVAAATNRVVAPTPGPGTPAERHADGLAL 62
Qv
            5 SVALVLLAAVLLQALLPASAEEGLVRIALKKRPIDENSRVAARLSG----EEGARRLGL 59
Db
          63 ALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGSSN----- 115
QУ
                60 RGANSLGGGGGEGDIVALKNYMNAQ----YFGEIGVGTPPQKFTVIFDTGSSNLWVPSAK 115
Db
         116 -- FAVAGTPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGFNTSF 173
Qу
              ]::| || :::||||: | ::| || || ||:
         116 CYFSIACFFHS----RYKSGQSSTYQKNGKPAAIQYGTGSIAGFFSEDSVTVGD----- 165
Db
         174 LVNIATIFESENFF----LPGI-----KWNGILGLAYATLAKPSSSLETFFDSLVTQANI 224
Qy
                  166 -----LVVKDQEFIEATKEPGLTFMVAKFDGILGLGFQEISVGDA------V 206
Db
         225 PNVFSMQMCG-AGLPVAGSGTN-----GGSLVLGGIEPSLYKGDIWYTPIKEEWYYQI 276
Qу
                                      || :| ||::|| |||: | |: :: |:|
             | : | | | | |
         207 PVWYKMVEQGLVSEPVFSFWFNRHSDEGEGGEIVFGGMDPSHYKGNHTYVPVSQKGYWQF 266
Db
         277 EILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPE---- 331
Qу
             1: : |||:: | : || |||||:|| | | : : | : : : : :
         267 EMGDVLIGGKTTGF-CA--SGCSAIADSGTSLLAGPTAIITEINEKIGATGVVSQECKTV 323
Db
                                                ----- 336
Qv
                                                          1:1
         324 VSQYGQQILDLLLAETQPSKICSQVGLCTFDGKHGVSAGIKSVVDDEAGESNGLQSGPMC 383
Db
```

```
337 -----FPKISIYLRD 364
Qy
                             : :|
                    1 : 1 | 1
         384 NACEMAVVWMQNQLAQNKTQDLILNYINQLCDKLPSPMGESSVDCGSLASMPEISFTIGA 443
Db
         365 ENSSRSFRITILPQLYIQPMMGAGLNYECY----RFGISPSTNAL-VIGATVMEGFYVIF 419
Qy
                                                      | | ::| | :: :|
                 : : |: || :| |
                                                1 1
         444 K-----KFALKPEEYIL-KVGEGAAAQCISGFTAMDIPPPRGPLWILGDVFMGAYHTVF 496
Dh
         420 DRAQKRVGFAAS 431
Qу
             1 : [[]]] [
         497 DYGKMRVGFAKS 508
Db
RESULT 12
PEPC RAT
                                 PRT;
                                        392 AA.
    PEPC RAT
                   STANDARD;
ID
AC
     P04073;
     01-NOV-1986 (Rel. 03, Created)
DT
     01-NOV-1986 (Rel. 03, Last sequence update)
DT
     28-FEB-2003 (Rel. 41, Last annotation update)
DT
     Gastricsin precursor (EC 3.4.23.3) (Pepsinogen C).
DΕ
GN
     Rattus norvegicus (Rat).
OS
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Mammalia; Eutheria; Rodentia; Sciurognathi; Muridae; Murinae; Rattus.
OC
     NCBI TaxID=10116;
OX
RN
     [1]
     SEQUENCE FROM N.A.
RP
RC
     STRAIN=Wistar;
     MEDLINE=89255508; PubMed=2722863;
RX
     Ishihara T., Ichihara Y., Hayano T., Katsura I., Sogawa K.,
RA
     Fujii-Kuriyama Y., Takahashi K.;
RA
     "Primary structure and transcriptional regulation of rat pepsinogen C
RT
     gene.";
RT
     J. Biol. Chem. 264:10193-10199(1989).
RL
RN
RP
     SEQUENCE FROM N.A.
RC
     STRAIN=Wistar;
     MEDLINE=87054020; PubMed=3780741;
RX
     Ichihara Y., Sogawa K., Morohashi K., Fujii-Kuriyama Y., Takahashi K.;
RA
     "Nucleotide sequence of a nearly full-length cDNA coding for
RT
     pepsinogen of rat gastric mucosa.";
RT
     Eur. J. Biochem. 161:7-12(1986).
RL
RN
     [3]
     SEQUENCE OF 16-112.
RP
RC
     STRAIN=Wistar;
     MEDLINE=84257697; PubMed=6743670;
RX
     Arai K.M., Muto N., Tani S., Akahane K.;
RA
     "The N-terminal sequence of rat pepsinogen.";
RT
     Biochim. Biophys. Acta 788:256-261(1984).
RL
     -!- CATALYTIC ACTIVITY: More restricted specificity than pepsin A, but
CC
         shows preferential cleavage at Tyr-|-Xaa bonds; high activity
CC
         towards hemoglobin as substrate.
CC
     -!- SIMILARITY: Belongs to peptidase family A1.
CC
     CC
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CC
CC
    EMBL; M25993; AAA41827.1; -.
DR
    EMBL; M25985; AAA41827.1; JOINED.
DR
    EMBL; M25986; AAA41827.1; JOINED.
DR
DR
    EMBL; M25987; AAA41827.1; JOINED.
    EMBL; M25988; AAA41827.1; JOINED.
DR
    EMBL; M25989; AAA41827.1; JOINED.
DR
    EMBL; M25990; AAA41827.1; JOINED.
DR
    EMBL; M25991; AAA41827.1; JOINED.
DR
    EMBL; M25992; AAA41827.1; JOINED.
DR
    EMBL; X04644; CAA28305.1; -.
DR
    PIR; A33510; A24608.
DR
    HSSP; P20142; 1AVF.
DR
    MEROPS; A01.003; -.
DR
    InterPro; IPR001969; Aspprotease_AS.
DR
    InterPro; IPR009007; Pept A acid.
    InterPro; IPR001461; Peptidase A1.
DR
    Pfam; PF00026; asp; 1.
DR
    PRINTS; PR00792; PEPSIN.
DR
    PROSITE; PS00141; ASP PROTEASE; 2.
DR
    Hydrolase; Aspartyl protease; Zymogen; Digestion; Signal.
KW
                1
                      16
FT
    SIGNAL
                17
                      62
                              ACTIVATION PEPTIDE.
    PROPEP
FT
                63
                     392
                              GASTRICSIN.
FT
    CHAIN
                     94
               94
    ACT SITE
FT
               280 280
    ACT SITE
FT
               107 112
                              BY SIMILARITY.
    DISULFID
FT
               270 275
                              BY SIMILARITY.
    DISULFID
FT
               314 347
                              BY SIMILARITY.
FT
    DISULFID
                              E \rightarrow Q (IN REF. 3).
               31
                     31
FT
    CONFLICT
               103
                              S \rightarrow A (IN REF. 3).
FT
                     103
    CONFLICT
                          s \rightarrow L \text{ (IN REF. 3).}
                     109
               109
FT
    CONFLICT
    SEQUENCE 392 AA; 42833 MW; 092A5EAF2783EDD1 CRC64;
SO
                       11.6%; Score 313; DB 1; Length 392;
  Query Match
  Best Local Similarity 29.5%; Pred. No. 2.2e-15;
  Matches 105; Conservative 56; Mismatches 139; Indels 56; Gaps
                                                                     16;
          92 YYLEMLIGTPPQKLQILVDTGSSNFAV-----AGTPHSYIDTYFDTERSSTYRSKGF 143
Qу
             76 YFGEISIGTPPQNFLVLFDTGSSNLWVSSVYCQSEACTTHA----RFNPSKSSTYYTEGQ 131
Db
         144 DVTVKYTQGSWTGFVGEDLVTIPKGFNTSFLVNIATIFESENFFLPG----IKWNGILG 198
Qy
               132 TFSLQYGTGSLTGFFGYDTLTV----QSIQVPNQEFGLSEN--EPGTNFVYAQFDGIMG 184
Db
         199 LAYATLAKPSSSLETFFDSLVTQANIPNVFSMQMCGAGLPVAGS--GTNGGSLVLGGIEP 256
QУ
             185 LAYPGLS--SGGATTALQGMLGE----GALSQPLFGVYL---GSQQGSNGGQIVFGGVDK 235
Db
         257 SLYKGDIWYTPIKEEWYYQIEILKLEIGGQSLNLDCREYNADKAIVDSGTTLLRLPQKVF 316
Qу
```

```
236 NLYTGEITWVPVTQELYWQITIDDFLIGDQASGW-CSSQGC-QGIVDTGTSLLVMPAQYL 293
Db
         317 DAVVEAVARASLIPEFSDGFWTGSQLACWTNSETPWSYFPKISIYLRDENSSRSFRITIL 376
Qy
                                                   | :| |
                                     ::|:|
                           1: : 1
         294 SELLQTIGAQE--GEYGEYF-----VSCDSVSS-----LPTLSFVL-----NGVQFPLS 335
Db
         377 PQLY-IQPMMGAGLNYECYRFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFAAS 431
QУ
                                          ::| : :| |||
                         : |
             336 PSSYIIQEDNFCMVGLESISLTSESGQPLWILGDVFLRSYYAIFDMGNNKVGLATS 391
Db
RESULT 13
PEPE CHICK
                                  PRT;
                                         383 AA.
     PEPE CHICK
                   STANDARD;
     P16476;
AC
     01-AUG-1990 (Rel. 15, Created)
DT
     01-AUG-1990 (Rel. 15, Last sequence update)
DT
     28-FEB-2003 (Rel. 41, Last annotation update)
DT
     Embryonic pepsinogen precursor (EC 3.4.23.-).
DE
     Gallus gallus (Chicken).
OS
     Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC
     Archosauria; Aves; Neognathae; Galliformes; Phasianidae; Phasianinae;
OC
OC
     Gallus.
     NCBI TaxID=9031;
OX
RN
     [1]
RP
     SEQUENCE FROM N.A.
     MEDLINE=88227903; PubMed=3131317;
RX
     Hayashi K., Agata K., Mochii M., Yasugi S., Eguchi G., Mizuno T.;
RA
     "Molecular cloning and the nucleotide sequence of cDNA for embryonic
RT
     chicken pepsinogen: phylogenetic relationship with prochymosin.";
RT
     J. Biochem. 103:290-296(1988).
RL
     -!- DEVELOPMENTAL STAGE: Specifically secreted during the embryonic
CC
         period in the chicken proventriculus (glandular stomach).
CC
     -!- SIMILARITY: Belongs to peptidase family A1.
CC
     CC
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     or send an email to license@isb-sib.ch).
CC
CC
DR
     EMBL; D00215; BAA00153.1; -.
DR
     PIR; A41443; A41443.
     HSSP; P00794; 4CMS.
DR
     MEROPS; A01.028; -.
DR
     InterPro; IPR001969; Aspprotease AS.
DR
     InterPro; IPR009007; Pept_A_acid.
DR
     InterPro; IPR001461; Peptidase_A1.
DR
     Pfam; PF00026; asp; 1.
DR
     PRINTS; PR00792; PEPSIN.
DR
     PROSITE; PS00141; ASP_PROTEASE; 2.
DR
     Hydrolase; Aspartyl protease; Digestion; Signal; Glycoprotein.
KW
                                  POTENTIAL.
     SIGNAL
                  1
                        16
FT
                                 EMBRYONIC PEPSINOGEN.
                  17
                        383
FT
     CHATN
                                 BY SIMILARITY.
     ACT SITE
                  94
                         94
FT
```

```
BY SIMILARITY.
    ACT SITE 276 276
FT
                           BY SIMILARITY.
             107
                   112
    DISULFID
FT
              267 271
310 344
                            BY SIMILARITY.
    DISULFID
FT
                            BY SIMILARITY.
   DISULFID 310 310 CARBOHYD 132 132 CARBOHYD 204 204 309
    DISULFID
FT
                            N-LINKED (GLCNAC. . .) (POTENTIAL).
FT
                            N-LINKED (GLCNAC. . .) (POTENTIAL).
FT
                            N-LINKED (GLCNAC. . .) (POTENTIAL).
    CARBOHYD 309 309
FT
                            N-LINKED (GLCNAC. . .) (POTENTIAL).
             350
                    350
FT
    CARBOHYD
             51 51
                            T \rightarrow S.
    VARIANT
FT
             383 AA; 41719 MW; 1642796871611F54 CRC64;
    SEOUENCE
SO
                     11.5%; Score 310; DB 1; Length 383;
 Query Match
 Best Local Similarity 26.8%; Pred. No. 3.5e-15;
 Matches 106; Conservative 63; Mismatches 136; Indels 90; Gaps 15;
         56 HA--DGLALALEPALASPAGAANFLAMVDNLQGDSGRGYYLEMLIGTPPQKLQILVDTGS 113
Qу
            55 HAFPDVLTVVTEPLL----NTLDM-----EYYGTISIGTPPQDFTVVFDTGS 97
Db
        114 SNFAVAG----TPHSYIDTYFDTERSSTYRSKGFDVTVKYTQGSWTGFVGEDLVTIPKGF 169
Qy
           98 SNLWVPSVSCTSPACQSHQMFNPSQSSTYKSTGQNLSIHYGTGDMEGTVGCDTVTVASLM 157
Db
        170 NTSFLVNIATIFESENFFLPGIKWNGILGLAYATLAKPSSSLETFFDSLVTQANI-PNVF 228
Qy
            :|: | ::| | || ||: :|::||||| ||:|| : : ||::| :: : |:|
        158 DTNQLFGLST-SEPGQFFV-YVKFDGILGLGYPSLA--ADGITPVFDNMVNESLLEQNLF 213
Db
        229 SMQMCGAGLPVAGSGTNGGSLVLGGIEPSLYKGDIWYTPIKEEWYYQIEILKLEIGGQSL 288
Qу
            214 SVYLSREPM-----GSMVVFGGIDESYFTGSINWIPVSYQGYWQISMDSIIVNKQEI 265
Db
        289 NLDCREYNADKAIVDSGTTLLRLPQKVFDAVVEAVARASLIPEFSDGFWTGSQLACWTNS 348
Qy
               : :||:|:|:|: | : : ||
        266 ACS----SGCQAIIDTGTSLVAGPASDINDIQSAVG------ANQ 300
Db
        349 ETPWSYFPKISIYLRDENSSRSFRITILPQLYIQPMMGAGLNY-----ECY 394
Qу
            301 NTYGEY----SVNCSHILAMPDVVF--VIG-GIQYPVPALAYTEQNGQGTCM 345
Db
        395 RFGISPSTNALVIGATVMEGFYVIFDRAQKRVGFA 429
Qу
              346 SSFQNSSADLWILGDVFIRVYYSIFDRANNRVGLA 380
Db
RESULT 14
CATD HUMAN
    CATD HUMAN STANDARD; PRT; 412 AA.
    P07339;
AC
    01-APR-1988 (Rel. 07, Created)
DT
    01-APR-1988 (Rel. 07, Last sequence update)
DT
    10-OCT-2003 (Rel. 42, Last annotation update)
DT
    Cathepsin D precursor (EC 3.4.23.5).
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    CTSD.
GN
    Homo sapiens (Human).
    Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
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    MEDLINE=85270436; PubMed=3927292;
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     Faust P.L., Kornfeld S., Chirgwin J.M.;
RA
     "Cloning and sequence analysis of cDNA for human cathepsin D.";
RT
     Proc. Natl. Acad. Sci. U.S.A. 82:4910-4914(1985).
RL
RN
     SEQUENCE FROM N.A.
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     MEDLINE=87231068; PubMed=3588310;
RX
     Westley B.R., May F.E.B.;
RA
     "Oestrogen regulates cathepsin D mRNA levels in oestrogen responsive
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     human breast cancer cells.";
RT
     Nucleic Acids Res. 15:3773-3786(1987).
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     SEQUENCE FROM N.A.
     MEDLINE=91299158; PubMed=2069717;
RX
     Redecker B., Heckendorf B., Grosch H.W., Mersmann G., Hasilik A.;
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     "Molecular organization of the human cathepsin D gene.";
RT
     DNA Cell Biol. 10:423-431(1991).
RL
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     Strausberg R.L., Feingold E.A., Grouse L.H., Derge J.G.,
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RA
     Altschul S.F., Zeeberg B., Buetow K.H., Schaefer C.F., Bhat N.K.,
RA
     Hopkins R.F., Jordan H., Moore T., Max S.I., Wang J., Hsieh F.,
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     Diatchenko L., Marusina K., Farmer A.A., Rubin G.M., Hong L.,
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     Stapleton M., Soares M.B., Bonaldo M.F., Casavant T.L., Scheetz T.E.,
RA
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RA
     Bosak S.A., McEwan P.J., McKernan K.J., Malek J.A., Gunaratne P.H.,
RA
     Richards S., Worley K.C., Hale S., Garcia A.M., Gay L.J., Hulyk S.W.,
RA
     Villalon D.K., Muzny D.M., Sodergren E.J., Lu X., Gibbs R.A.,
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     Fahey J., Helton E., Ketteman M., Madan A., Rodrigues S., Sanchez A.,
RA
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RA
     Blakesley R.W., Touchman J.W., Green E.D., Dickson M.C.,
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     Rodriguez A.C., Grimwood J., Schmutz J., Myers R.M.,
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     Butterfield Y.S.N., Krzywinski M.I., Skalska U., Smailus D.E.,
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     "Generation and initial analysis of more than 15,000 full-length
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     Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903(2002).
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     MEDLINE=94085791; PubMed=8262386;
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     May F.E., Smith D.J., Westley B.R.;
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     "The human cathepsin D-encoding gene is transcribed from an estrogen-
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     regulated and a constitutive start point.";
RT
     Gene 134:277-282(1993).
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     Augereau P., Miralles F., Cavailles V., Gaudelet C., Parker M.,
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      "Characterization of the proximal estrogen-responsive element of
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Mol. Endocrinol. 8:693-703(1994).
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     Hochstrasser D.F., Frutiger S., Paquet N., Bairoch A., Ravier F.,
RA
     Pasquali C., Sanchez J.-C., Tissot J.-D., Bjellqvist B., Vargas R.,
RA
     Appel R.D., Hughes G.J.;
RA
     Submitted (JUN-1992) to Swiss-Prot.
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     MEDLINE=22660472; PubMed=12754519;
RX
     Zhang H., Li X.-J., Martin D.B., Aebersold R.;
RA
     "Identification and quantification of N-linked glycoproteins using
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RT
     Nat. Biotechnol. 21:660-666(2003).
RL
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     Papassotiropoulos A., Bagli M., Kurz A., Kornhuber J., Forstl H.,
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     Maier W., Pauls J., Lautenschlager N., Heun R.;
RA
     "A genetic variation of cathepsin D is a major risk factor for
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RT
     Ann. Neurol. 47:399-403(2000).
RL
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     X-RAY CRYSTALLOGRAPHY (3 ANGSTROMS).
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     TISSUE=Spleen;
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     MEDLINE=93223670; PubMed=8467789;
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     Metcalf P., Fusek M.;
RA
     "Two crystal structures for cathepsin D: the lysosomal targeting
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     signal and active site.";
RT
     EMBO J. 12:1293-1302(1993).
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RN
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     Baldwin E.T., Bhat T.N., Gulnik S., Hosur M.V., Sowder R.C. II,
RA
     Cachau R.E., Collins J., Silva A.M., Erickson J.W.;
RA
     "Crystal structures of native and inhibited forms of human cathepsin
RT
     D: implications for lysosomal targeting and drug design.";
RT
     Proc. Natl. Acad. Sci. U.S.A. 90:6796-6800(1993).
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         Involved in the pathogenesis of several diseases such as breast
CC
         cancer and possibly Alzheimer's disease.
CC
     -!- CATALYTIC ACTIVITY: Specificity similar to, but narrower than,
CC
         that of pepsin A. Does not cleave the 4-Gln-|-His-5 bond in B
CC
         chain of insulin.
CC
     -!- SUBUNIT: Consists of a light chain and a heavy chain.
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CC
         risk for developing AD than noncarriers.
CC
     -!- SIMILARITY: Belongs to peptidase family A1.
CC
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     the European Bioinformatics Institute. There are no restrictions on its
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     or send an email to license@isb-sib.ch).
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Qу
             62 VTEGPI--PEVLKNYM-----DAQYYGEIGIGTPPQCFTVVFDTGSSNLWVPSIH 109
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Db
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Qy
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DT
     15-MAR-2004 (Rel. 43, Last annotation update)
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     Diedrich J.F., Staskus K.A., Retzel E.F., Haase A.T.;
RA
     "Nucleotide sequence of a cDNA encoding mouse cathepsin D.";
RT
     Nucleic Acids Res. 18:7184-7184(1990).
RL
RN
RP
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     MEDLINE=90326544; PubMed=2374732;
     Grusby M.J., Mitchell S.C., Glimcher L.H.;
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RT
     "Molecular cloning of mouse cathepsin D.";
RL
     Nucleic Acids Res. 18:4008-4008(1990).
RN
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     Hetman M., Perschl A., Saftig P., von Figura K., Peters C.;
RA
     "Mouse cathepsin D gene: molecular organization, characterization of
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RA
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     Fahey J., Helton E., Ketteman M., Madan A., Rodrigues S., Sanchez A.,
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RA
     Schnerch A., Schein J.E., Jones S.J.M., Marra M.A.;
RA
     "Generation and initial analysis of more than 15,000 full-length
RT
     human and mouse cDNA sequences.";
RT
     Proc. Natl. Acad. Sci. U.S.A. 99:16899-16903(2002).
RL
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CC
    -!- FUNCTION: Acid protease active in intracellular protein breakdown.
    -!- CATALYTIC ACTIVITY: Specificity similar to, but narrower than,
CC
        that of pepsin A. Does not cleave the 4-Gln-|-His-5 bond in B
CC
CC
        chain of insulin.
CC
    -!- SUBUNIT: Consists of a light chain and a heavy chain.
CC
    -!- SUBCELLULAR LOCATION: Lysosomal.
CC
    -!- SIMILARITY: Belongs to peptidase family Al.
    _____
CC
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    the European Bioinformatics Institute. There are no restrictions on its
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    use by non-profit institutions as long as its content is in no way
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    or send an email to license@isb-sib.ch).
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FT
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FT
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FT
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               110 117
                              BY SIMILARITY.
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               284
FT
    DISULFID
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FT
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Qy
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